

WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification 6:

A01N 43/54 // (A01N 43/54, 61:00, 55:00, 47:38, 43:84, 43:653, 43:54, 37:50)

(11) International Publication Number:

WO 98/33381

A1

(43) International Publication Date:

6 August 1998 (06.08.98)

(21) International Application Number:

PCT/US98/01381

(22) International Filing Date:

26 January 1998 (26.01.98)

(30) Priority Data:

60/036,063

30 January 1997 (30.01.97)

(71) Applicant (for all designated States except US): E.I. DU PONT DE NEMOURS AND COMPANY [US/US]; 1007 Market Street, Wilmington, DE 19898 (US).

(72) Inventors: and

- (75) Inventors/Applicants (for US only): CROMPTON, Martina, Cajnar [US/US]; 6178 Telegraph Road, Elkton, MD 21921 (US). FOOR, Stephen, Ray [US/US]; 508 Hemingway Drive, Hockessin, DE 19707 (US). JULIS, Anthony, Jay [US/US]; 2706 Doris Drive, Wilmington, DE 19803 (US).
- (74) Agent: HEISER, David, E.; E.I. du Pont de Nemours and Company, Legal Patent Records Center, 1007 Market Street, Wilmington, DE 19898 (US).

(81) Designated States: AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GW, HU, ID, IL, IS, JP, KG, KP, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA, US, UZ, VN, YU, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).

Published

With international search report.

Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.

(54) Title: FUNGICIDAL MIXTURES

(57) Abstract

Advantageous combinations are disclosed comprising (1) at least one compound selected from the quinazolinones of Formula (I), N-oxides and agriculturally suitable salts thereof and (2) at least one compound selected from: (a) compounds acting at the bci complex of the fungal mitochondrial respiratory electron transfer site and (b) compounds that control fungal disease by inhibiting sterol biosynthesis. D is O or S and R1-R4 are as defined in the disclosure. Included are fungicidal

$$R^3$$
 N R^1 OR^2

compositions comprising fungicidally effective amounts of the compound combinations of the invention and a method for controlling plant diseases caused by fungal plant pathogens comprising applying to the plant or portion thereof, or to the plant seed or seedling, a fungicidally effective amount of the compound combinations of the invention.

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

	•						• •	
AL	Albania	ES	Spain	LS	Lesotho	SÍ	Slovenia	
AM	Armenia	FI	Finland .	LT	Lithuania	· sk	Slovakia	
AΤ	Austria	FR	France	LU	Luxembourg	SN	Senegal	
AU	Australia	GA	Gabon	· LV	Latvia	SZ	Swaziland	
AZ	Azerbaijan	GB	United Kingdom	MC	Monaco	TD	Chad	
BA	Bosnia and Herzegovina	GE	Georgia	MD	Republic of Moldova	TG	Togo	
BB	Barbados	GH	Ghana	MG	Madagascar	TJ	Tajikistan	
BE	Belgium	GN	Guinea	MK	The former Yugoslav	TM	Turkmenistan	
BF	Burkina Faso	GR	Greece		Republic of Macedonia	TR	Turkey	
BG	Bulgaria	HU	Hungary	ML	Mali	ΤŤ	Trinidad and Tobago	
Bj	Benin	IE	Ireland	MN.	Mongolia	UA	Ukraine	
BR	Brazil	IL.	Israel	MR	Mauritania	UG	Uganda .	
BY	Belarus	IS	Iceland	MW	Malawi	US	United States of America	
CA	Canada	IT	Italy	MX	Mexico	UZ	Uzbekistan	
CF	Central African Republic	JP	Japan	NE	Niger	VN	Viet Nam	
CG	Congo	KE	Kenya	. NL	Netherlands	YU	Yugoslavia	
CH	Switzerland	КG	Kyrgyzstan	NO	Norway	zw	Zimbabwe	
CI	Côte d'Ivoire	KP	Democratic People's	NZ	New Zealand			
CM	Cameroon		Republic of Korea	PL	Poland			
CN	China	KR	Republic of Korea	PT	Portugal			
CU	Cuba	KZ	Kazakstan	RO	Romania			
CZ	Czech Republic	LC	Saint Lucia	RU	Russian Federation			
DE	Germany	LI	Liechtenstein	SD	Sudan			
DK	_ Denmark	LK	Sri Lanka	SE	Sweden			
EE	Estonia	LR	Liberia	SG	Singapore			

WO 98/33381 PCT/US98/01381

1

TITLE FUNGICIDAL MIXTURES BACKGROUND OF THE INVENTION

Fungicides that effectively control plant diseases are in constant demand by growers. Plant diseases are highly destructive, difficult to control and quickly develop resistance to commercial fungicides. Combinations of pesticides are often used to facilitate disease control, to broaden spectrum of control and to retard resistance development. It is recognized in the art that the advantages of particular pesticide combinations can often vary, depending on such factors as the particular plant and plant disease to be treated, and the treatment conditions. Accordingly, there is an ongoing search for advantageous combinations of pesticides.

5

10

International Patent Application WO 94/26722 discloses certain quinazolinone compounds as fungicides (e.g., 6-bromo-3-propyl-2-propyloxy-4(3H)-quinazolinone, 6,8-diiodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and 6-iodo-3-propyl-2propyloxy-4(3H)-quinazolinone). International Patent Application WO 95/14009 15 discloses certain triazolone fungicides (e.g., 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]-methyl]phenyl]-3H-1,2,4triazol-3-one). Kresoxim-methyl (BAS 490F) is a fungicide under consideration for the control of plant diseases, especially fungal diseases of wheat, grapes and fruits. (See E. Ammermann, G. Lorenz, K. Schelberger, B. Wenderoth, H. Sauter and C. Rentzea 20 "BAS 490F - A Broad-Spectrum Fungicide with a New Mode of Action" in Brighton Crop Protection Conference - Pests and Diseases - 1992, 1, 403-410.) Azoxystrobin (ICIA5504) is a fungicide under consideration for the control of plant diseases on a wide range of crops. (See J. R. Godwin, V. M. Anthony, J. M. Clough and C. R. A. Godfrey "ICIA5504: A Novel, Broad Spectrum, Systemic β-Methoxyacrylate Fungicide" in 25 Brighton Crop Protection Conference - Pests and Diseases - 1992, 1, 435-442.) European Patent Application EP-A-398,692 discloses (E)-2-(methoxyimino)-N-methyl-2-(2-phenoxyphenyl)acetamide also known as SSF 126. This fungicide is under consideration for the control of plant disease, especially of rice. EP-A-68813 discloses 30 certain triazole compounds as fungicides (e.g., flusilazole). EP-A-40345 discloses

certain triazole compounds as fungicides (e.g., tebuconazole).

SUMMARY OF THE INVENTION

This invention is directed to fungicidal combinations (e.g., mixtures) comprising (1) at least one compound selected from the quinazolinones of Formula I (including all geometric and stereoisomers), N-oxides, and agriculturally suitable salts thereof,

R³ 5 D 3 R¹ 7 R⁴ 8 N 1 OR²

wherein

5

10

20

25

D is O or S;

 R^1 is C_1 - C_{10} alkyl; C_3 - C_5 cycloalkyl; C_3 - C_{10} alkenyl; C_3 - C_{10} alkynyl;

C₁-C₁₀ haloalkyl; C₃-C₁₀ haloalkenyl; C₃-C₁₀ haloalkynyl;

C₂-C₁₀ alkoxyalkyl; C₂-C₁₀ alkylthioalkyl; C₂-C₁₀ alkylsulfinylalkyl;

C2-C10 alkylsulfonylalkyl; C4-C10 cycloalkylalkyl;

 C_4 - C_{10} alkenyloxyalkyl; C_4 - C_{10} alkynyloxyalkyl; C_4 - C_{10}

(cycloalkyl)oxyalkyl; C₄-C₁₀ alkenylthioalkyl; C₄-C₁₀ alkynylthioalkyl;

15 C₄-C₁₀ (cycloalkyl)thioalkyl; C₂-C₁₀ haloalkoxyalkyl;

C₄-C₁₀ haloalkenyloxyalkyl; C₄-C₁₀ haloalkynyloxyalkyl;

 C_4 - C_{10} alkoxyalkenyl; C_4 - C_{10} alkoxyalkynyl; C_4 - C_{10} alkylthioalkenyl;

 C_4 - C_{10} alkylthioalkynyl; C_4 - C_{10} trialkylsilylalkyl; C_1 - C_{10} alkyl substituted

with NR⁵R⁶, nitro, cyano or phenyl optionally substituted with R⁸, R⁹ and

 R^{10} ; C_1 - C_{10} alkoxy; C_1 - C_{10} haloalkoxy; C_1 - C_{10} alkylthio;

C₁-C₁₀ haloalkylthio; or pyridinyl, furanyl, thienyl, naphthalenyl,

benzofuranyl, benzothienyl or quinolinyl each optionally substituted with R⁸, R⁹ and R¹⁰:

 R^2 is C_1 - C_{10} alkyl; C_3 - C_7 cycloalkyl; C_3 - C_{10} alkenyl; C_3 - C_{10} alkynyl;

C₁-C₁₀ haloalkyl; C₃-C₁₀ haloalkenyl; C₃-C₁₀ haloalkynyl;

 C_2 - C_{10} alkoxyalkyl; C_2 - C_{10} alkylthioalkyl; C_2 - C_{10} alkylsulfinylalkyl;

 C_2 - C_{10} alkylsulfonylalkyl; C_4 - C_{10} cycloalkylalkyl;

C₄-C₁₀ alkenyloxyalkyl; C₄-C₁₀ alkynyloxyalkyl; C₄-C₁₀

(cycloalkyl)oxyalkyl; C₄-C₁₀ alkenylthioalkyl; C₄-C₁₀ alkynylthioalkyl;

30 C₄-C₁₀ (cycloalkyl)thioalkyl; C₂-C₁₀ haloalkoxyalkyl;

 C_4 - C_{10} haloalkenyloxyalkyl; C_4 - C_{10} haloalkynyloxyalkyl;

 C_4 - C_{10} alkoxyalkenyl; C_4 - C_{10} alkoxyalkynyl; C_4 - C_{10} alkylthioalkenyl;

C₄-C₁₀ alkylthioalkynyl; C₄-C₁₀ trialkylsilylalkyl; C₂-C₁₀ cyanoalkyl;

15

20

25

30

35

 C_1 - C_{10} nitroalkyl; C_1 - C_{10} alkyl substituted with CO_2R^5 , NR^5R^6 , or phenyl optionally substituted with R^7 , R^9 and R^{10} ; $N=CR^5R^5$; or $N=CR^5R^6$; or

 R^1 and R^2 are taken together to form -CH₂(CH₂)_qCH₂-;

5 q is 0, 1, 2, 3 or 4;

R³ is halogen, C₁-C₈ alkyl, C₃-C₈ cycloalkyl, C₂-C₈ alkenyl, C₂-C₈ alkynyl, C₁-C₈ haloalkyl, C₃-C₈ haloalkenyl, C₃-C₈ haloalkynyl, C₁-C₈ alkoxy, C₁-C₈ haloalkoxy, C₃-C₈ alkenyloxy, C₃-C₈ alkynyloxy, C₁-C₈ alkylthio, C₃-C₈ alkenylthio, C₃-C₈ alkynylthio, C₁-C₈ alkylsulfinyl, C₁-C₈ alkylsulfonyl, C₂-C₈ alkoxyalkyl, C₂-C₈ alkylsulfonylalkyl, C₄-C₈ cycloalkylalkyl, C₃-C₈ trialkylsilyl, NR⁵R⁶, C₅-C₈ trialkylsilylalkynyl or phenyl optionally substituted with at least one R⁷;

 R^4 is hydrogen, halogen, C_1 - C_4 alkyl, C_1 - C_4 haloalkyl, C_1 - C_4 alkoxy or C_1 - C_4 haloalkoxy;

each R⁵ is independently hydrogen, C₁-C₄ alkyl or phenyl optionally substituted with at least one R⁷;

each R⁶ is independently hydrogen, C₁-C₈ alkyl or phenyl optionally substituted with at least one R⁷; or

each R⁷ is independently halogen, C₁-C₄ alkyl, C₁-C₄ alkoxy, C₁-C₄ haloalkyl, nitro or cyano;

R⁸ is C₁-C₆ alkyl, C₁-C₆ alkoxy, C₁-C₆ haloalkyl, halogen, C₂-C₈ alkynyl, C₁-C₆ alkylthio, phenyl or phenoxy each optionally substituted with at least one R⁷, cyano, nitro, C₁-C₆ haloalkoxy, C₁-C₆ haloalkylthio, C₂-C₆ alkenyl, C₂-C₆ haloalkenyl, acetyl, CO₂Me or N(C₁-C₂ alkyl)₂;

each R⁹ is independently methyl, ethyl, methoxy, methylthio, halogen or trifluoromethyl; and

each R¹⁰ is independently halogen;

and (2) at least one compound selected from (a) compounds acting at the bc_1 complex of the fungal mitochondrial respiratory electron transfer site and (b) compounds that control fungal disease by inhibiting the sterol biosynthesis pathway. This invention provides agricultural compositions containing these combinations and the use of the combinations as fungicides. Advantageous compositions include those where component (1) and component (2) are present in a fungicidally effective amount and the

10

15

20

25

mole ratio of component (1) to component (2) is from about 30:1 to 1:30. Advantageous methods include those where component (1) and component (2) are added in amounts sufficient to provide a fungicidal effectiveness greater than the sum of the fungicidal effectivenesses provided by those amounts of said components taken independently.

DETAILS OF THE INVENTION

Combinations of fungicides are used in accordance with this invention to facilitate disease control and to retard resistance development. Suitable compositions and methods are provided.

For example, this invention also provides methods for controlling plant diseases caused by fungal plant pathogens comprising applying to the plant or portion thereof to be protected, or to the plant seed or seedling to be protected one of the following:

- A) an effective amount of a fungicidal composition comprising component (1), component (2), and at least one of a surfactant, a solid diluent or a liquid diluent;
- B) (i) an effective amount of a first composition comprising component (1), and at least one of a surfactant, solid or liquid diluent; and (ii) an effective amount of a second composition comprising component (2), and at least one of a surfactant, a solid diluent or a liquid diluent; said first and second compositions applied sequentially in any order; or
- C) an effective amount of a physical mixture of the first and second compositions as defined in B above.

The mole ratio of the compound(s) of component (1) to the compound(s) of component (2) applied is normally from about 30:1 to 1:30, and the compound(s) of component (1) and the compound(s) of component (2) are normally applied in amounts effective to provide control of the fungal disease which is greater than the additive control of that fungal disease provided by the compound(s) of component (1) and the compound(s) of component (2) individually.

Preferred compositions for reasons of ease of synthesis or greater fungicidal activity are:

Preferred 1. A fungicidal composition comprising a fungicidally effective amount of (1) at least one quinazolinone of Formula I (including all geometric and stereoisomers), N-oxides, and agriculturally suitable salts thereof, wherein

 R^1 is C_1 - C_6 alkyl;

R² is C₁-C₆ alkyl;

R³ is halogen and fixed at position 6 of the quinazolinone; and R⁴ is hydrogen or halogen and fixed at position 8 of the quinazolinone; and (2) at least one compound selected from

(a) compounds of Formula II (including all geometric and stereoisomers), N-oxides, and agriculturally suitable salts thereof,

II

5

wherein

Q is

$$X \longrightarrow A \longrightarrow N$$
 or $MeO \longrightarrow N$

Q-1

Q-2

WlMe

10

15

E is selected from:

- i) 1,2-phenylene optionally substituted with R^{13} or both R^{13} and R^{14} :
- ii) a naphthalene ring, provided that when G and Y are attached to the same ring, then G and Y are attached to adjacent ring members, the naphthalene ring optionally substituted with R¹³ or both R¹³ and R¹⁴; and
- iii) a ring system selected from 5 to 12-membered monocyclic and fused bicyclic aromatic heterocyclic ring systems, each heterocyclic ring system containing 1 to 6 heteroatoms independently selected from the group nitrogen, oxygen, and sulfur, provided that each heterocyclic ring system contains no more than 4 nitrogens, no more than 2 oxygens, and no more than 2 sulfurs, each fused bicyclic ring system optionally containing one nonaromatic ring that optionally includes one or two J¹ as ring members and optionally includes one or two ring members independently selected from C(=O) and S(O)₂, provided that G is attached to an aromatic ring, and when G and Y are attached to the same ring, then G and Y are attached to adjacent ring members, each aromatic

20

25

heterocyclic ring system optionally substituted with R^{13} or both R^{13} and R^{14} ;

A is O, S, N, NR¹⁵ or CR²⁴;

G is C or N; provided that when G is C, then A is O, S or NR¹⁵ and the floating double bond is attached to G; and when G is N, then A is N or CR²⁴ and the floating double bond is attached to A;

W is O, S, NH, $N(C_1-C_6 \text{ alkyl})$ or $NO(C_1-C_6 \text{ alkyl})$;

W1 is O or NH;

X is H, OR¹¹, S(O)_mR¹¹, halogen, C₁-C₆ alkyl, C₁-C₆ haloalkyl, C₃-C₆ cycloalkyl, cyano, NH₂, NHR¹¹, N(C₁-C₆ alkyl)R¹¹, NH(C₁-C₆ alkoxy) or N(C₁-C₆ alkoxy)R¹¹;

X1 is CH or N;

R¹¹ is C₁-C₆ alkyl, C₁-C₆ haloalkyl, C₂-C₆ alkenyl, C₂-C₆ haloalkenyl, C₂-C₆ alkynyl, C₂-C₆ haloalkynyl, C₃-C₆ cycloalkyl, C₂-C₄ alkylcarbonyl or C₂-C₄ alkoxycarbonyl;

R¹² is H, C₁-C₆ alkyl, C₁-C₆ haloalkyl, C₂-C₆ alkenyl, C₂-C₆ haloalkenyl, C₂-C₆ alkynyl, C₂-C₆ haloalkynyl,

C₃-C₆ cycloalkyl, C₂-C₄ alkylcarbonyl, C₂-C₄ alkoxycarbonyl, hydroxy, C₁-C₂ alkoxy or acetyloxy;

R¹³ and R¹⁴ are each independently halogen; cyano; nitro; hydroxy; C₁-C₆ alkyl; C₁-C₆ haloalkyl; C₂-C₆ alkenyl; C₂-C₆ haloalkenyl;

C₂-C₆ alkynyl; C₂-C₆ haloalkynyl; C₁-C₆ alkoxy;

C₁-C₆ haloalkoxy; C₂-C₆ alkenyloxy; C₂-C₆ alkynyloxy;

 $C_1\hbox{-} C_6 \ alkylthio; \ C_1\hbox{-} C_6 \ alkylsulfinyl; \ C_1\hbox{-} C_6 \ alkylsulfonyl; \ formyl;$

C₂-C₆ alkylcarbonyl; C₂-C₆ alkoxycarbonyl; NH₂C(O);

 $(C_1-C_4 \text{ alkyl})\text{NHC}(O); (C_1-C_4 \text{ alkyl})_2\text{NC}(O); \text{Si}(R^{35})_3; \text{Ge}(R^{35})_3;$

 $(R^{35})_3Si-C=C-$; or phenyl, phenylethynyl, benzoyl, or

phenylsulfonyl each substituted with R¹⁸ and optionally substituted with one or more R²⁰; or

when E is 1,2-phenylene and R¹³ and R¹⁴ are attached to adjacent atoms, R¹³ and R¹⁴ can be taken together as C₃-C₅ alkylene, C₃-C₅ haloalkylene, C₃-C₅ alkenylene or C₃-C₅ haloalkenylene

 R^{15} is H, C_1 - C_6 alkyl, C_1 - C_6 haloalkyl, C_2 - C_6 alkenyl,

each optionally substituted with 1-2 C₁-C₃ alkyl;

C2-C6 haloalkenyl, C2-C6 alkynyl, C2-C6 haloalkynyl,

 C_3 - C_6 cycloalkyl, C_2 - C_4 alkylcarbonyl or C_2 - C_4 alkoxycarbonyl;

Y is -O-, -S(O)_n-, -NR²⁵-, -C(=O)-, -CH(OR²⁵)-, -CHR¹⁶-, -CHR¹⁶CHR¹⁶-, -CR¹⁶=CR¹⁶-, -C≡C-, -CHR²⁵O-, -OCHR²⁵-,

.

5

10

15

20

25

30

35

```
-CHR^{25}S(O)_{n}, -S(O)_{n}CHR^{25}, -CHR^{25}O-N=C(R^{17}),
                                   -(R^{17})C=N-OCH(R^{25})-, -C(R^{17})=N-O-, -O-N=C(R^{17})-,
                                   -CHR^{25}OC(=O)N(R^{25})-, -CHR^{25}OC(=S)N(R^{25})-,
                                   -CHR<sup>25</sup>OC(=O)O-, -CHR<sup>25</sup>OC(=S)O-, -CHR<sup>25</sup>OC(=O)S-,
                                   -CHR^{25}OC(=S)S-, -CHR^{25}SC(=O)N(R^{25})-,
 5
                                   -CHR<sup>25</sup>SC(=S)N(R<sup>25</sup>)-, -CHR<sup>25</sup>SC(=O)O-, -CHR<sup>25</sup>SC(=S)O-,
                                   -CHR<sup>25</sup>SC(=O)S-, -CHR<sup>25</sup>SC(=S)S-, -CHR<sup>25</sup>SC(=NR<sup>25</sup>)S-,
                                   -CHR^{25}N(R^{25})C(=O)N(R^{25})-, -CHR^{25}O-N(R^{25})C(=O)N(R^{25})-,
                                   -CHR<sup>25</sup>O-N(R<sup>25</sup>)C(=S)N(R<sup>25</sup>)-, -CHR<sup>25</sup>O-N=C(R<sup>17</sup>)NR<sup>25</sup>-,
10
                                   -CHR^{25}O-N=C(R^{17})OCH_{2-}, -CHR^{25}O-N=C(R^{17})-N=N_{-}
                                   -CHR^{25}O-N=C(R^{17})-C(=O)-
                                   -CHR^{25}O-N=C(R^{17})-C(=N-A^2-Z^1)-A^1
                                   -CHR^{25}O-N=C(R^{17})-C(R^{17})=N-A^2-A^3
                                   -CHR^{25}O-N=C(-C(R^{17})=N-A^2-Z^1)-, -CHR^{25}O-N=C(R^{17})-CH_2O-,
                                   -CHR<sup>25</sup>O-N=C(R<sup>17</sup>)-CH<sub>2</sub>S-, -O-CH<sub>2</sub>CH<sub>2</sub>O-N=C(R<sup>17</sup>)-,
15
                                   -CHR^{25}O-C(R^{25})=C(R^{17})-, -CHR^{25}O-C(R^{17})=N-,
                                   -CHR<sup>25</sup>S-C(R<sup>17</sup>)=N-, -C(R<sup>17</sup>)=N-NR<sup>25</sup>-, -CH=N-N=C(R<sup>17</sup>)-,
                                   -CHR^{25}N(R^{25})-N=C(R^{17})-, -CHR^{25}N(COCH_3)-N=C(R^{17})-,
                                   -OC(=S)NR^{25}C(=O)-, -CHR^{16}-C(=W^2)-A^1-,
                                   -CHR16CHR16-C(=W2)-A1-, -CR16=CR16-C(=W2)-A1-,
20
                                   -C = C - C(=W^2) - A^1, -N = CR^{16} - C(=W^2) - A^1 or a direct bond; and
                                   the directionality of the Y linkage is defined such that the moiety
                                   depicted on the left side of the linkage is bonded to E and the
                                   moiety on the right side of the linkage is bonded to Z;
                           Z^1 is H or -A<sup>3</sup>-Z;
25
                           W^2 is O or S:
                           A<sup>1</sup> is O, S, NR<sup>25</sup> or a direct bond;
                           A<sup>2</sup> is O, NR<sup>25</sup> or a direct bond;
                           A^3 is -C(=O)-, -S(O)2- or a direct bond;
                           each R16 is independently H, 1-2 CH3, C2-C3 alkyl, C1-C3 alkoxy,
30
                                   C<sub>3</sub>-C<sub>6</sub> cycloalkyl, formylamino, C<sub>2</sub>-C<sub>4</sub> alkylcarbonylamino,
                                   C2-C4 alkoxycarbonylamino, NH2C(O)NH,
                                   (C<sub>1</sub>-C<sub>3</sub> alkyl)NHC(O)NH, (C<sub>1</sub>-C<sub>3</sub> alkyl)<sub>2</sub>NC(O)NH,
                                   N(C<sub>1</sub>-C<sub>3</sub> alkyl)<sub>2</sub>, piperidinyl, morpholinyl, 1-2 halogen, cyano or
35
                                   nitro:
                           each R<sup>17</sup> is independently H, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> haloalkyl,
                                   C1-C6 alkoxy, C1-C6 haloalkoxy, C1-C6 alkylthio,
                                   C<sub>1</sub>-C<sub>6</sub> alkylsulfinyl, C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl, C<sub>1</sub>-C<sub>6</sub> haloalkylthio,
```

C₁-C₆ haloalkylsulfinyl, C₁-C₆ haloalkylsulfonyl, C₂-C₆ alkenyl, C₂-C₆ haloalkenyl, C₂-C₆ alkynyl, C₂-C₆ haloalkynyl, C₃-C₆ cycloalkyl, C₂-C₄ alkylcarbonyl, C₂-C₄ alkoxycarbonyl, halogen, cyano, nitro, hydroxy, amino, NH(C₁-C₆ alkyl), N(C₁-C₆ alkyl)₂ or morpholinyl; each Z is independently selected from:

i) C₁-C₁₀ alkyl, C₂-C₁₀ alkenyl and C₂-C₁₀ alkynyl each substituted with R¹⁹ and optionally substituted with one or more R²⁰.

10

5

ii) C₃-C₈ cycloalkyl, C₃-C₈ cycloalkenyl and phenyl each substituted with R¹⁹ and optionally substituted with one or more R²⁰;

15

iii) a ring system selected from 3 to 14-membered monocyclic, fused bicyclic and fused tricyclic nonaromatic heterocyclic ring systems and 5 to 14-membered monocyclic, fused bicyclic and fused tricyclic aromatic heterocyclic ring systems, each heterocyclic ring system containing 1 to 6 heteroatoms independently selected from the group nitrogen, oxygen, and sulfur,

20

provided that each heterocyclic ring system contains no more than 4 nitrogens, no more than 2 oxygens, and no more than 2 sulfurs, each nonaromatic or aromatic heterocyclic ring system substituted with R¹⁹ and optionally substituted with one or more R²⁰;

25.

iv) a multicyclic ring system selected from 8 to 14-membered fused-bicyclic and fused-tricyclic ring systems which are an

aromatic carbocyclic ring system, a nonaromatic carbocyclic ring system, or a ring system containing one or two nonaromatic rings that each include one or two J¹ as ring members and one or two ring members independently selected from C(=O) and S(O)₂, and

any remaining rings as aromatic carbocyclic rings, each multicyclic ring system substituted with R^{19} and optionally substituted with

30

v) adamantyl substituted with R^{19} and optionally substituted with one or more R^{20} ;

35

each J¹ is independently selected from the group -CHR²³-, -NR²³-, -O-and -S(O)_p-;

R¹⁸ is H, 1-2 halogen, C₁-C₆ alkyl, C₁-C₆ haloalkyl, C₁-C₆ alkoxy, C₁-C₆ haloalkoxy, C₂-C₆ alkenyl, C₂-C₆ haloalkenyl, C₂-C₆ alkynyl, C₁-C₆ alkylthio, C₁-C₆ haloalkylthio,

one or more R²⁰; and

	·
	C ₁ -C ₆ alkylsulfinyl, C ₁ -C ₆ alkylsulfonyl, C ₃ -C ₆ cycloalkyl,
	C_3 - C_6 alkenyloxy, $CO_2(C_1$ - C_6 alkyl), $NH(C_1$ - C_6 alkyl),
	$N(C_1-C_6 \text{ alkyl})_2$, cyano, nitro, $SiR^{29}R^{30}R^{31}$ or $GeR^{29}R^{30}R^{31}$;
•	R^{19} is H; 1-2 halogen; C_1 - C_6 alkyl; C_1 - C_6 haloalkyl; C_1 - C_6 alkoxy;
5	C ₁ -C ₆ haloalkoxy; C ₂ -C ₆ alkenyl; C ₂ -C ₆ haloalkenyl;
	C ₂ -C ₆ alkynyl; C ₁ -C ₆ alkylthio; C ₁ -C ₆ haloalkylthio;
	C ₁ -C ₆ alkylsulfinyl; C ₁ -C ₆ alkylsulfonyl; C ₃ -C ₆ cycloalkyl;
	C_3 - C_6 alkenyloxy; $CO_2(C_1$ - C_6 alkyl); $NH(C_1$ - C_6 alkyl);
	$N(C_1-C_6 \text{ alkyl})_2$; $-C(R^{28})=NOR^{27}$; cyano; nitro; SF ₅ ;
10	SiR ³² R ³³ R ³⁴ ; or GeR ³² R ³³ R ³⁴ ; or R ¹⁹ is phenyl, benzyl, benzoyl,
	phenoxy, pyridinyl, pyridinyloxy, thienyl, thienyloxy, furanyl,
	pyrimidinyl or pyrimidinyloxy each optionally substituted with R ²¹
	or both R^{21} and R^{22} ;
	each R^{20} is independently halogen, C_1 - C_4 alkyl, C_1 - C_4 haloalkyl,
15	C ₁ -C ₄ alkoxy, nitro or cyano; or
	when R ¹⁹ and an R ²⁰ are attached to adjacent atoms on Z, R ¹⁹ and said
	adjacently attached R ²⁰ can be taken together as -OCH ₂ O- or
	-OCH ₂ CH ₂ O-; each CH ₂ group of said taken together R ¹⁹ and R ²⁰
	optionally substituted with 1-2 halogen; or
20	when Y and an R ²⁰ are attached to adjacent atoms on Z and Y is
	-CHR ²⁵ O-N=C(R ¹⁷)-, -O-N=C(R ¹⁷)-, -O-CH ₂ CH ₂ O-N=C(R ¹⁷)-,
	-CHR ²⁵ O-C(R ²⁵)=C(R ¹⁷)-, -CH=N-N=C(R ¹⁷)-,
	-CHR ²⁵ N(R ²⁵)-N=C(R ¹⁷)- or -CHR ²⁵ N(COCH ₃)-N=C(R ¹⁷)-, R ¹⁷
	and said adjacently attached R ²⁰ can be taken together as
25	-(CH ₂) _r -J- such that J is attached to Z;
	J is -CH ₂ -, -CH ₂ CH ₂ -, -OCH ₂ -, -CH ₂ O-, -SCH ₂ -, -CH ₂ S-, -N(R ²⁶)CH ₂ -
	or -CH ₂ N(R ²⁶)-; each CH ₂ group of said J optionally substituted
•	with 1 to 2 CH ₃ ;
	R ²¹ and R ²² are each independently 1-2 halogen; C ₁ -C ₄ alkyl;
30	C ₁ -C ₄ haloalkyl; C ₂ -C ₆ alkenyl; C ₂ -C ₆ haloalkenyl; C ₂ -C ₆
	alkynyl; C ₂ -C ₆ haloalkynyl; C ₂ -C ₆ alkoxyalkyl; C ₂ -C ₆
	alkylthioalkyl; C ₃ -C ₆ alkoxyalkynyl;
	C ₇ -C ₁₀ tetrahydropyranyloxyalkynyl; benzyloxymethyl; C ₁ -C ₄
	alkoxy; C ₁ -C ₄ haloalkoxy; C ₃ -C ₆ alkenyloxy; C ₃ -C ₆
35	haloalkenyloxy; C ₃ -C ₆ alkynyloxy; C ₃ -C ₆ haloalkynyloxy; C ₂ -C ₆
	alkoxyalkoxy; C5-C9 trialkylsilylalkoxyalkoxy;
	C ₂ -C ₆ alkylthioalkoxy; C ₁ -C ₄ alkylthio; C ₁ -C ₄ haloalkylthio;
	C_1 - C_4 alkylsulfinyl; C_1 - C_4 haloalkylsulfinyl; C_1 - C_4 alkylsulfonyl;

C₁-C₄ haloalkylsulfonyl; C₃-C₆ alkenylthio; C₃-C₆ haloalkenylthio; C₂-C₆ alkylthioalkylthio; nitro; cyano; thiocyanato; hydroxy; N(R³⁶)₂; SF₅; Si(R³⁵)₃; Ge(R³⁵)₃; $(R^{35})_3Si-C=C-$; $OSi(R^{35})_3$; $OGe(R^{35})_3$; $C(=O)R^{36}$; $C(=S)R^{36}$; $C(=O)OR^{36}$; $C(=S)OR^{36}$; $C(=O)SR^{36}$; $C(=S)SR^{36}$; $C(=O)N(R^{36})_2$; 5 $C(=S)N(R^{36})_2$; $OC(=O)R^{36}$; $OC(=S)R^{36}$; $SC(=O)R^{36}$; $SC(=S)R^{36}$; $N(R^{36})C(=O)R^{36}$; $N(R^{36})C(=S)R^{36}$; $OC(=O)OR^{37}$; $OC(=O)SR^{37}$; $OC(=O)N(R^{36})_2$; $SC(=O)OR^{37}$; $SC(=O)SR^{37}$; $S(O)_2OR^{36}$; $S(O)_2N(R^{36})_2$; $OS(O)_2R^{37}$; $N(R^{36})S(O)_2R^{37}$; or phenyl, phenoxy, 10 benzyl, benzyloxy, phenylsulfonyl, phenylethynyl or pyridinylethynyl, each optionally substituted with halogen, C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy, C₁-C₄ haloalkoxy, nitro or cyano: each R²³ is independently H; C₁-C₆ alkyl; C₁-C₆ haloalkyl; or phenyl 15 optionally substituted with halogen, C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy, C₁-C₄ haloalkoxy, nitro or cyano; R²⁴ is H, halogen, C₁-C₆ alkyl, C₁-C₆ haloalkyl, C₂-C₆ alkenyl, C2-C6 haloalkenyl, C2-C6 alkynyl, C2-C6 haloalkynyl or C₃-C₆ cycloalkyl; 20 each R²⁵ is independently H; C₁-C₃ alkyl; C₃-C₆ cycloalkyl; or phenyl or benzyl, each optionally substituted on the phenyl ring with halogen, C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy, C₁-C₄ haloalkoxy, nitro or cyano; or when Y is -CHR²⁵N(R²⁵)C(=O)N(R²⁵)-, the two R²⁵ attached to 25 nitrogen atoms on said group can be taken together as -(CH₂)_s-; or when Y is -CHR²⁵O-N=C(R¹⁷)NR²⁵-, R¹⁷ and the adjacently attached R²⁵ can be taken together as -CH₂-(CH₂)_S-, -O-(CH₂)_S-, -S-(CH₂)_S- or -N(C₁-C₃ alkyl)-(CH₂)_S-; with the directionality of said linkage defined such that the moiety depicted on the left side 30 of the linkage is bonded to the carbon and the moiety on the right side of the linkage is bonded to the nitrogen; R²⁶, R²⁷ and R²⁸ are each independently H; C₁-C₃ alkyl; C₃-C₆ cycloalkyl; or phenyl optionally substituted with halogen,

35

 R^{29} , R^{30} , R^{31} , R^{32} , R^{33} and R^{34} are each independently C_1 - C_6 alkyl, C_2 - C_6 alkenyl, C_1 - C_4 alkoxy or phenyl;

nitro or cyano;

C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy, C₁-C₄ haloalkoxy,

each R35 is independently C1-C4 alkyl, C1-C4 haloalkyl, C2-C4 alkenyl, C₁-C₄ alkoxy or phenyl; each R³⁶ is independently H; C₁-C₆ alkyl; C₁-C₆ haloalkyl; C2-C6 alkenyl; C2-C6 haloalkenyl; C2-C6 alkynyl; C2-C6 haloalkynyl; C3-C6 cycloalkyl; or phenyl or benzyl, each optionally substituted on the phenyl ring with halogen, C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy, C₁-C₄ haloalkoxy, nitro or cyano; each R³⁷ is independently C₁-C₆ alkyl; C₁-C₆ haloalkyl; C₂-C₆ alkenyl; C₂-C₆ haloalkenyl; C₂-C₆ alkynyl; C₂-C₆ haloalkynyl; C₃-C₆ cycloalkyl; or phenyl or benzyl, each optionally substituted 10 on the phenyl ring with halogen, C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy, C₁-C₄ haloalkoxy, nitro or cyano; m, n and p are each independently 0, 1 or 2; r is 0 or 1; and 15 s is 2 or 3; and (b) bromuconazole, cyproconazole, difenoconazole, diniconazole, epoxiconazole, fenarimol, fenbuconazole, fenpropidin, fenpropimorph, fluquinconazole, flusilazole, flutriafol, hexaconazole, ipconazole, metconazole, penconazole, 20 propiconazole, tebuconazole, tetraconazole, triadimefon, triadimenol, tridemorph, triticonazole and uniconazole. Preferred 2. The fungicidal composition of Preferred 1 comprising a fungicidally effective amount of (1) at least one compound selected from the group consisting of 25 6-bromo-3-propyl-2-propyloxy-4(3H)-quinazolinone, 6,8-diiodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; and (2) at least one compound selected from the group consisting of 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-30 (trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl] phenyl]-3H-1,2,4-triazol-3-one, kresoxim-methyl, azoxystrobin, (E)-2-(methoxyimino)-N-methyl-2-(2phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and tebuconazole. Preferred 3. The fungicidal composition of Preferred 2 comprising a fungicidally 35 effective amount of (1) 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone (sometimes referred to hereafter as the Formula Ia compound) and (2) 2,4-

dihydro-5-methoxy-2-methyl-4-[2-[[[[1-[3-

PCT/US98/01381

WO 98/33381

5

30

35

12

(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4triazol-3-one (sometimes referred to hereafter as the Formula IIa compound).

- Preferred 4. The fungicidal composition of Preferred 2 comprising a fungicidally effective amount of (1) 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and (2) flusilazole.
- Preferred 5. The fungicidal composition of Preferred 2 comprising a fungicidally effective amount of (1) 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and (2) tebuconazole.
- Preferred 6. The fungicidal composition of Preferred 2 comprising a fungicidally 10 effective amount of (1) 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone, and (2) both 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4triazol-3-one and flusilazole.
- In the above recitations, the term "alkyl", used either alone or in compound words · 15 such as "alkylthio" or "haloalkyl" includes straight-chain or branched alkyl, such as, methyl, ethyl, n-propyl, i-propyl, or the different butyl, pentyl, hexyl, heptyl, octyl, nonyl and decyl isomers. The term "1-2 CH3" indicates that the substituent can be methyl (i.e., Me) or, when there is a hydrogen attached to the same atom, the substituent and said hydrogen can both be methyl. "Alkenyl" includes straight-chain or branched 20 alkenes such as ethenyl, 1-propenyl, 2-propenyl, and the different butenyl, pentenyl, hexenyl, heptenyl, octenyl, nonenyl and decenyl isomers. "Alkenyl" also includes polyenes such as 1,2-propadienyl and 2,4-hexadienyl. "Alkynyl" includes straight-chain or branched alkynes such as ethynyl, 1-propynyl, 2-propynyl and the different butynyl, pentynyl, hexynyl, heptynyl, octynyl, nonynyl, and decynyl isomers. "Alkynyl" can 25 also include moieties comprised of multiple triple bonds such as 2,5-hexadiynyl.
 - "Alkylene" denotes a straight-chain or branched alkanediyl. Examples of "alkylene" include CH2CH2CH2, CH2CH(CH3) and the different butylene and pentylene isomers. "Haloalkylene" denotes a halogen substituted alkylene. Examples of "haloalkylene" include CH₂CH(CF₃), CH₂CF₂CH₂ and CH₂CH(CCl₃). "Alkenylene" denotes a straight-chain or branched alkenediyl containing one olefinic bond. Examples of "alkenylene" include CH2CH=CH and CH=C(CH3) and the different butenylene and pentenylene isomers. "Haloalkenylene" denotes a halogen substituted alkenylene. Examples of "haloalkenylene" include CH2CCl=CCl and $CH=C(CF_3)$.
 - "Alkenyloxyalkyl" denotes alkenyl substitution on oxygen which in turn is substituted on alkyl. Examples "alkenyloxyalkyl" include CH2=CHCH2OCH2 and CH₃CH=CHCH₂OCH₂CH₂. "Alkynyloxyalkyl" denotes alkynyl substitution on

15

20

25

oxygen which in turn is substituted on alkyl. Examples of "alkynyloxyalkyl" include $CH = CCH_2OCH_2$ and $CH_3C = CCH_2OCH_2CH_2$.

"Alkoxy" includes, for example, methoxy, ethoxy, propyloxy, 1-methylethoxy and the different butoxy, pentyloxy, hexyloxy, heptyloxy, octyloxy, nonyloxy, and decyloxy isomers. "Alkoxyalkyl" denotes alkoxy substitution on alkyl. Examples of "alkoxyalkyl" include CH₃OCH₂, CH₃OCH₂CH₂, CH₃CH₂OCH₂, CH₃CH₂CH₂CH₂OCH₂ and CH₃CH₂OCH₂CH₂. "Alkoxyalkoxy" denotes alkoxy substitution on alkoxy. Examples of "alkoxyalkoxy" include CH₃OCH₂O, (CH₃)₃COCH₂O and CH₃OCH₂CH₂O. "Alkenyloxy" includes straight-chain or branched alkenyloxy moieties. Examples of "alkenyloxy" include H₂C=CHCH₂O, (CH₃)₂C=CHCH₂O, (CH₃)CH=CHCH₂O, (CH₃)CH=C(CH₃)CH₂O and CH₂=CHCH₂CH₂O. "Alkynyloxy" includes straight-chain or branched alkynyloxy moieties. Examples of "alkynyloxy" include HC≡CCH2O, CH3C≡CCH2O and CH₃C=CCH₂CH₂O. "Alkoxyalkenyl" denotes alkoxy substitution of alkenyl. "Alkoxyalkenyl" includes straight-chain or branched alkoxyalkenyl moieties. Examples of "alkoxyalkenyl" include (CH₃)₂CHOCH=CH and CH₃OCH₂CH=CH. "Alkoxyalkynyl" denotes alkoxy substitution of alkynyl. "Alkoxyalkynyl" includes straight-chain or branched alkoxyalkynyl moieties. Examples of "alkoxyalkynyl" include $(CH_3)_2CHOCH_2C\equiv C$ and $CH_3OCH_2C\equiv C$.

- "Alkylthioalkoxy" denotes alkylthio substitution on alkoxy. Examples of "alkylthioalkoxy" include CH₃SCH₂O and CH₃CH₂SCH₂CH₂O. "Alkylthioalkylthio" denotes alkylthio substitution on alkylthio. Examples of "alkylthioalkylthio" include CH₃SCH₂S and CH₃SCH₂CH₂S. "Alkylsulfinyl" includes both enantiomers of an alkylsulfinyl group. Examples of "alkylsulfinyl" include CH₃S(O), CH₃CH₂S(O),
- CH₃CH₂CH₂S(O), (CH₃)₂CHS(O) and the different butylsulfinyl, pentylsulfinyl, hexylsulfinyl, heptylsulfinyl and octylsulfinyl isomers. "Alkylsulfinylalkyl" denotes alkylsulfinyl substitution on alkyl. Examples of "alkylsulfinylalkyl" include CH₃S(O)CH₂, CH₃S(O)CH₂CH₂, CH₃CH₂S(O)CH₂, CH₃CH₂CH₂CH₂CH₂CH₂CH₂CH₂ and CH₃CH₂S(O)CH₂CH₂. Examples of "alkylsulfonyl" include CH₃S(O)₂,
- CH₃CH₂S(O)₂, CH₃CH₂CH₂S(O)₂, (CH₃)₂CHS(O)₂ and the different butylsulfonyl, pentylsulfonyl, hexylsulfonyl, heptylsulfonyl and octylsulfonyl isomers.

 "Alkylsulfonylalkyl" denotes alkylsulfonyl substitution on alkyl. Examples of "alkylsulfonylalkyl" include CH₃S(O)₂CH₂, CH₃S(O)₂CH₂CH₂, CH₃CH₂S(O)₂CH₂,

 $CH_3CH_2CH_2CH_2S(O)_2CH_2 \ and \ CH_3CH_2S(O)_2CH_2CH_2. \ "Alkylthioalkenyl" \ denotes alkylthio substitution on alkenyl. Examples of "alkylthioalkenyl" include <math display="block">CH_3SCH_2CH=CH \ and \ CH_3CH_2SCH=CH. \ "Alkylthioalkynyl" \ denotes alkylthio substitution on alkynyl. Examples of "alkylthioalkynyl" include <math>CH_3SCH_2C\equiv C$ and $CH_3CH_2SCH_2CH_2C\equiv C. \ "Alkenylthio" includes straight-chain and branched alkenylthio moieties. Examples of "alkenylthio" include <math>CH_2=CHCH_2S$ and $CH_2=CHCH_2CH_2S. \ "Alkenylthioalkyl" \ denotes alkenylthio substitution on alkyl. Examples of "alkenylthioalkyl" include <math>CH_2=CHCH_2SCH_2$ and $CH_2=CHCH_2CH_2SCH_2. \ "Alkynylthio" includes straight-chain and branched$ alkynylthio moieties. Examples of "alkynylthio" include $CH\equiv CCH_2S$ and $CH\equiv CCH_2CH_2S. \ "Alkynylthioalkyl" \ denotes alkynylthio substitution on alkyl. Examples of "alkynylthioalkyl" include <math>CH\equiv CCH_2SCH_2$ and $CH\equiv CCH_2CH_2SCH_2$.

"Cycloalkyl" includes, for example, cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl, cycloheptyl and cyclooctyl. "(Cycloalkyl)oxyalkyl" denotes cycloalkyl substituted on oxygen which in turn is substituted on alkyl. Examples of "(cycloalkyl)oxyalkyl" include (cyclopentyloxy)methyl and (cyclohexyloxy)methyl. "(Cycloalkyl)thioalkyl" denotes cycloalkyl substituted on sulfur which in turn is substituted on alkyl. Examples of "(cycloalkyl)thioalkyl" include (cyclopentylthio)methyl and (cyclohexylthio)methyl. "Cycloalkenyl" includes groups such as cyclopentenyl and cyclohexenyl as well as groups with more than one double bond such as 1,3- and 1,4-cyclohexadienyl. Examples of "cycloalkylalkyl" include cyclopropylmethyl, cyclopentylethyl, and other cycloalkyl moieties bonded to straight-chain or branched alkyl groups.

15

20

25

30

35

"Cyanoalkyl" denotes an alkyl group substituted with one cyano group. Examples of "cyanoalkyl" include NCCH₂, NCCH₂CH₂ and CH₃CH(CN)CH₂. "Nitroalkyl" denotes an alkyl group substituted with one nitro group. Examples of "nitroalkyl" include NO₂CH₂ and CH₃CH(NO₂)CH₂.

"Tetrahydropyranyloxyalkynyl" denotes a tetrahydropyranyl group on oxygen which in turn is substituted on an alkynyl group. An example of "tetrahydropyranyloxyalkynyl" is 2-[(tetrahydro-2*H*-pyranyl)oxy]ethynyl.

The term "aromatic ring system" denotes fully unsaturated carbocycles and heterocycles in which the polycyclic ring system is aromatic (where aromatic indicates that the Hückel rule is satisfied for the ring system). The term "aromatic carbocyclic ring system" includes fully aromatic carbocycles and carbocycles in which at least one ring of a polycyclic ring system is aromatic (where aromatic indicates that the Hückel rule is satisfied). The term "nonaromatic carbocyclic ring system" denotes fully saturated carbocycles as well as partially or fully unsaturated carbocycles where the Hückel rule is not satisfied by any of the rings in the ring system. The term "aromatic

10

15

20

25

30

35

heterocyclic ring system" includes fully aromatic heterocycles and heterocycles in which at least one ring of a polycyclic ring system is aromatic (where aromatic indicates that the Hückel rule is satisfied). Examples of "aromatic heterocyclic ring systems" include furanyl, furazanyl, thienyl, pyrrolyl, pyrazolyl, oxazolyl, oxadiazolyl, imidazolyl, isoxazolyl, thiazolyl, thiadiazolyl, isothiazolyl, pyridyl, pyrimidinyl, pyridazinyl, pyrazinyl and triazinyl with said ring attached through any available carbon or nitrogen. For example, when the aromatic heterocyclic ring is furanyl, it can be 2-furanyl or 3-furanyl, for pyrrolyl, the aromatic heterocyclic ring is 1-pyrrolyl, 2-pyrrolyl or 3-pyrrolyl, for pyridyl, the aromatic ring is 2-pyridyl, 3-pyridyl or 4-pyridyl and similarly for other aromatic heterocyclic rings. The term "nonaromatic heterocyclic ring system" denotes fully saturated heterocycles as well as partially or fully unsaturated heterocycles where the Hückel rule is not satisfied by any of the rings in the ring system. The heterocyclic ring systems can be attached through any available carbon or nitrogen by replacement of a hydrogen on said carbon or nitrogen.

One skilled in the art will appreciate that not all nitrogen containing heterocycles can form N-oxides since the nitrogen requires an available lone pair for oxidation to the oxide; one skilled in the art will recognize those nitrogen containing heterocycles which can form N-oxides. One skilled in the art will also recognize that tertiary amines can form N-oxides. Synthetic methods for the preparation of N-oxides of heterocycles and tertiary amines are very well known by one skilled in the art including the oxidation of heterocycles and tertiary amines with peroxy acids such as peracetic and m-chloroperbenzoic acid (MCPBA), hydrogen peroxide, alkyl hydroperoxides such as t-butyl hydroperoxide, sodium perborate, and dioxiranes such as dimethydioxirane. These methods for the preparation of N-oxides have been extensively described and reviewed in the literature, see for example: T. L. Gilchrist in Comprehensive Organic Synthesis, vol. 7, pp 748-750, S. V. Ley, Ed., Pergamon Press; M. Tisler and B. Stanovnik in Comprehensive Heterocyclic Chemistry, vol. 3, pp 18-20, A. J. Boulton and A. McKillop, Eds., Pergamon Press; M. R. Grimmett and B. R. T. Keene in Advances in Heterocyclic Chemistry, vol. 43, pp 149-161, A. R. Katritzky, Ed., Academic Press; M. Tisler and B. Stanovnik in Advances in Heterocyclic Chemistry, vol. 9, pp 285-291, A. R. Katritzky and A. J. Boulton, Eds., Academic Press; and G. W. H. Cheeseman and E. S. G. Werstiuk in Advances in Heterocyclic Chemistry, vol. 22, pp 390-392, A. R. Katritzky and A. J. Boulton, Eds., Academic Press.

The term "halogen", either alone or in compound words such as "haloalkyl", includes fluorine, chlorine, bromine or iodine. The term "1-2 halogen" indicates that one or two of the available positions for that substituent may be halogen which are independently selected. Further, when used in compound words such as "haloalkyl", said alkyl may be partially or fully substituted with halogen atoms which may be the

same or different. Examples of "haloalkyl" include F₃C, ClCH₂, CF₃CH₂ and CF₃CCl₂. The terms "haloalkenyl", "haloalkenyloxy", "haloalkenylthio", "haloalkenyloxyalkyl", "haloalkynyl", "haloalkynyloxy", "haloalkynyloxyalkyl", "haloalkoxy", "haloalkoxyalkyl", "haloalkylthio", "haloalkylsulfinyl",

- "haloalkylsulfonyl", and the like, are defined analogously to the term "haloalkyl".

 Examples of "haloalkenyl" include (Cl)₂C=CHCH₂ and CF₃CH₂CH=CHCH₂.

 Examples of "haloalkenyloxy" include (Cl)₂C=CHCH₂O and CF₃CH₂CH=CHCH₂O.

 Examples of "haloalkenylthio" include (Cl)₂C=CHCH₂S and CF₃CH₂CH=CHCH₂S.

 Examples of "haloalkenyloxyalkyl" include (Cl)₂C=CHCH₂OCH₂ and
- 10 CF₃CH₂CH=CHCH₂OCH₂. Examples of "haloalkynyl" include HC=CCHCl, CF₃C=C, CCl₃C=C and FCH₂C=CCH₂. Examples of "haloalkynyloxy" include CF₃C=CCH₂O, CCl₃C=CCH₂O and FCH₂C=CCH₂O. Examples of "haloalkynyloxyalkyl" include CCl₃C=CCH₂OCH₂ and FCH₂C=CCH₂CH₂OCH₂. Examples of "haloalkoxy" include CF₃O, CCl₃CH₂O, HCF₂CH₂CH₂O and CF₃CH₂O. Examples of "haloalkoxyalkyl"
- include CF₃OCH₂, CCl₃CH₂OCH₂CH₂, HCF₂ CH₂CH₂OCH₂ and CF₃ CH₂OCH₂. Examples of "haloalkylthio" include CCl₃S, CF₃S, CCl₃CH₂S and ClCH₂CH₂CH₂S. Examples of "haloalkylsulfinyl" include CF₃S(O), CCl₃S(O), CF₃CH₂S(O) and CF₃CF₂S(O). Examples of "haloalkylsulfonyl" include CF₃S(O)₂, CCl₃S(O)₂, CF₃CH₂S(O)₂ and CF₃CF₂S(O)₂.

"Alkylcarbonyl" denotes alkyl substituted carbonyl. Examples of "alkylcarbonyl" include CH₃C(=O) and (CH₃)₂CHC(=O). "Alkoxycarbonyl denotes alkoxy substituted carbonyl. Examples of "alkoxycarbonyl" include CH₃OC(=O) and (CH₃)₂CHOC(=O). "Alkylcarbonylamino" denotes alkylcarbonyl substituted on nitrogen. Examples of "alkylcarbonylamino" include CH₃C(=O)NH and CH₃CH₂C(=O)NH.

"Alkoxycarbonylamino" denotes alkoxycarbonyl substituted on nitrogen. Examples of "alkoxycarbonylamino" include CH₃OC(=0)NH and CH₃CH₂OC(=0)NH.

Examples of "trialkylsilyl" include (CH₃)₃Si and (CH₃)₃CSi(CH₃)₂. "Trialkylsilylalkyl" denotes trialkylsilyl substitution on alkyl. Examples of "trialkylsilylalkyl" include (CH₃)₃SiCH₂, and (CH₃)₃SiCH₂CH₂.

- "Trialkylsilylalkynyl" denotes trialkylsilyl substitution on alkynyl. Examples of "trialkylsilylalkynyl" include (CH₃)₃SiC≡C and (CH₃CH₂)SiCH₂C≡C.
 "Trialkylsilylalkoxyalkoxy" denotes trialkylsilyl substitution on alkoxy substituted in turn on alkoxy. Examples of "trialkylsilylalkoxyalkoxy" include (CH₃)₃SiCH₂OCH₂O and (CH₃)₃SiCH₂CCH₂OCH₂O.
- The total number of carbon atoms in a substituent group is indicated by the "C_i-C_j" prefix where i and j are numbers from 1 to 11. For example, C₁-C₃ alkylsulfonyl designates methylsulfonyl through propylsulfonyl; C₂ alkoxyalkyl designates CH₃OCH₂; C₃ alkoxyalkyl designates, for example, CH₃CH(OCH₃),

PCT/US98/01381

17

CH₃OCH₂CH₂ or CH₃CH₂OCH₂; and C₄ alkoxyalkyl designates the various isomers of an alkyl group substituted with an alkoxy group containing a total of four carbon atoms, examples including CH₃CH₂CH₂OCH₂ and CH₃CH₂OCH₂CH₂. Examples of "alkylcarbonyl" include C(O)CH₃, C(O)CH₂CH₂CH₃ and C(O)CH(CH₃)₂. Examples of "alkoxycarbonyl" include CH₃OC(=O), CH₃CH₂OC(=O), CH₃CH₂CH₂OC(=O), (CH₃)₂CHOC(=O) and the different butoxy- or pentoxycarbonyl isomers. In the above recitations, when a compound of Formula I is comprised of one or more heterocyclic rings, all substituents are attached to these rings through any available carbon or nitrogen by replacement of a hydrogen on said carbon or nitrogen.

When a compound is substituted with a substituent bearing a subscript that indicates the number of said substituents can exceed 1, said substituents (when they exceed 1) are independently selected from the group of defined substituents. Further, when the subscript indicates a range, e.g., (R)_{i-j}, then the number of substituents may be selected from the integers between i and j inclusive.

10

15

20

25

30

35

When a group contains a substituent which can be hydrogen, for example R4, then, when this substituent is taken as hydrogen, it is recognized that this is equivalent to said group being unsubstituted. When a group is optionally substituted with a substituent, for example with R7, then, when the group is not substituted with that substituent, it is recognized that this is equivalent to said group having a hydrogen substituent.

Compounds used in this invention often can exist as one or more stereoisomers. The various stereoisomers include enantiomers, diastereomers, atropisomers and geometric isomers. One skilled in the art will appreciate that one stereoisomer may be more active and/or may exhibit beneficial effects when enriched relative to the other stereoisomer(s) or when separated from the other stereoisomer(s). (See, e.g., U.S. Provisional Patent Application Serial No. 60/057917 filed September 4, 1997, which is hereby incorporated by reference in its entirety.) Additionally, the skilled artisan knows how to separate, enrich, and/or to selectively prepare said stereoisomers. Accordingly, the present invention comprises compounds selected from Formula I, N-oxides and agriculturally suitable salts thereof. The compounds of the invention may be present as a mixture of stereoisomers, individual stereoisomers, or as an optically active form.

The salts of the compounds which may used in the invention include acid-addition salts with inorganic or organic acids such as hydrobromic, hydrochloric, nitric, phosphoric, sulfuric, acetic, butyric, fumaric, lactic, maleic, malonic, oxalic, propionic, salicylic, tartaric, 4-toluenesulfonic or valeric acids. The salts of the compounds which may be used in the invention also include those formed with organic bases (e.g., pyridine, ammonia, or triethylamine) or inorganic bases (e.g., hydrides, hydroxides, or

PCT/US98/01381

carbonates of sodium, potassium, lithium, calcium, magnesium or barium) when the compound contains an acidic group such as a carboxylic acid or phenol.

The fungicidal compositions of this invention, in addition to comprising fungicidally effective amounts of the mixtures of the invention, also optionally comprise at least one of a surfactant, a solid diluent or a liquid diluent. The preferred compositions of the present invention are those which comprise the above preferred component (1) and component (2) compounds.

This invention also relates to a method for controlling plant diseases caused by fungal plant pathogens comprising applying to the plant or portion thereof, or to the plant seed or seedling, a fungicidally effective amount of the compositions of the invention (e.g., as a composition described herein). The preferred methods of use include those involving the above preferred compositions.

The bc₁ Complex Fungicides

5

10

25

Compounds of Formula II are known to have a fungicidal mode of action which inhibits the bc₁ complex. Methyl (E)-2-[[6-(2-cyanophenoxy)-4-pyrimidinyl]oxy]-α-(methoxyimino)benzeneacetate is described as a bc₁ complex inhibitor in Biochemical Society Transactions 1993, 22, 68S. Methyl (E)-α-(methoxyimino)-2-[(2-methylphenoxy)methyl]benzeneacetate is described as a bc₁ complex inhibitor in Biochemical Society Transactions 1993, 22, 64S. (E)-2-[(2,5-

20 Dimethylphenoxy)methyl]- α -(methoxyimino)-N-methylbenzeneacetamide is described as a bc_1 complex inhibitor in *Biochemistry and Cell Biology* 1995, 85(3), 306-311.

The bc_1 complex is sometimes referred to by other names in the biochemical literature, including complex III of the electron transfer chain, and ubihydroquinone:cytochrome c oxidoreductase. It is uniquely identified by the Enzyme Commission number EC1.10.2.2. The bc_1 complex is described in, for example, *J. Biol. Chem.* 1989, 264, 14543-38; *Methods Enzymol.* 1986, 126, 253-71; and references cited therein.

The Sterol Biosynthesis Inhibitor Fungicides

The class of sterol biosynthesis inhibitors includes DMI and non-DMI compounds, that control fungi by inhibiting enzymes in the sterol biosynthesis pathway. DMI fungicides have a common site of action within the fungal sterol biosynthesis pathway; that is, an inhibition of demethylation at position 14 of lanosterol or 24-methylene dihydrolanosterol, which are precursors to sterols in fungi. Compounds acting at this site are often referred to as demethylase inhibitors, DMI fungicides, or DMIs. The demethylase enzyme is sometimes referred to by other names in the biochemical literature, including cytochrome P-450 (14DM). The demethylase enzyme is described in, for example, *J. Biol. Chem.* 1992, 267, 13175-79 and references cited therein. DMI fungicides fall into several classes: triazoles, imidazoles, pyrimidines,

piperazines and pyridines. The triazoles includes bromuconazole, cyproconazole, difenoconazole, diniconazole, epoxiconazole, fenbuconazole, fluquinconazole, flusilazole, flutriafol, hexaconazole, ipconazole, metconazole, penconazole, propiconazole, tebuconazole, tetraconazole, triadimefon, triadimenol, triticonazole and uniconazole. The imidazoles include clotrimazole, econazole, imazalil, isoconazole, miconazole and prochloraz. The pyrimidines include fenarimol, nuarimol and triarimol. The piperazines include triforine. The pyridines include buthiobate and pyrifenox. Biochemical investigations have shown that all of the above mentioned fungicides are DMI fungicides as described by K. H. Kuck, et al. in *Modern Selective Fungicides - Properties, Applications and Mechanisms of Action*, Lyr, H., Ed.; Gustav Fischer Verlag: New York, 1995, 205-258.

The DMI fungicides have been grouped together to distinguish them from other sterol biosynthesis inhibitors, such as, the morpholine and piperidine fungicides. The morpholines and piperidines are also sterol biosynthesis inhibitors but have been shown to inhibit later steps in the sterol biosynthesis pathway. The morpholines include aldimorph, dodemorph, fenpropimorph, tridemorph and trimorphamide. The piperidines include fenpropidin. Biochemical investigations have shown that all of the above mentioned morpholine and piperidine fungicides are sterol biosynthesis inhibitor fungicides as described by K. H. Kuck, et al. in *Modern Selective Fungicides - Properties, Applications and Mechanisms of Action*, Lyr, H., Ed.; Gustav Fischer Verlag: New York, 1995, 185-204.

Synergistic Effects

10

15

20

25

30

35

Fungicides that effectively control plant fungi, particularly wheat powdery mildew (Erysiphe graminis) and wheat foot rot (Pseudocercosporella herpotrichoides), are in constant demand by growers. Combinations of fungicides are often used to facilitate disease control and to retard resistance development. Mixtures of fungicides may provide significantly better disease control than could be predicted based on the activity of the individual components. This synergism has been described as "the cooperative action of two components of a mixture, such that the total effect is greater or more prolonged than the sum of the effects of the two (or more) taken independently" (see Tames, P. M. L., Neth. J. Plant Pathology, (1964), 70, 73-80). It has been demonstrated that compositions containing compounds of Formula Ia and Formula IIa; Formula Ia and flusilazole; Formula Ia and tebuconazole; and Formula Ia, Formula IIa and flusilazole exhibit synergistic effects.

The presence of a synergistic effect between two active ingredients is established with the aid of the Colby equation (see Colby, S. R. In *Calculating Synergistic and Antagonistic Responses of Herbicide Combinations*, Weeds, (1967), 15, 20-22):

10

15

20

25

30

35

$$p = A + B - \left[\frac{A \times B}{100} \right]$$

Using the methods of the Colby, the presence of a synergistic interaction between two active ingredients is established by first calculating the predicted activity, p, of the mixture based on activities of the two components applied alone. If p is lower than the experimentally established effect, synergism has occurred. In the equation above, A is the fungicidal activity in percentage control of one component applied alone at rate x. The B term is the fungicidal activity in percentage control of the second component applied at rate y. The equation estimates p, the fungicidal activity of the mixture of A at rate x with B at rate y if their effects are strictly additive and no interaction has occurred.

The Colby equation for a three way mixture is

$$p = A + B + C + \frac{A \times B \times C}{10000} - \frac{(A \times B) + (A \times C) + (B \times C)}{100}$$

In this application, fungicidal activities provided by compositions of Formula Ia, Formula IIa, flusilazole and tebuconazole alone are compared with that of compositions of the compounds of Formula Ia and Formula IIa together, compositions of the compounds of Formula Ia and flusilazole together, compositions of the compounds of Formula Ia and tebuconazole together, and compositions of the compounds of Formula Ia, Formula IIa and flusilazole together. Based on the description of synergism developed by Colby, compositions of the present invention are considered to be synergistically useful. Accordingly, this invention provides an improved method of combating fungi, such as wheat powdery mildew (Erysiphe graminis), wheat leaf rust (Puccinia recondita), wheat foot rot (Pseudocercosporella herpotrichoides) and/or wheat glume blotch (Septoria nodorum) in crops, especially cereals.

Compositions are provided in accordance with this invention which comprise proportions of component (1) and component (2) which are especially useful for controlling particular fungal diseases. For example, the compositions of this invention include those wherein the mole ratio of component (1) to component (2) is from about 30:1 to 1:30. These compositions are considered especially useful for controlling wheat powdery mildew (*Erysiphe graminis*), wheat leaf rust (*Puccinia recondita*), wheat foot rot (*Pseudocercosporella herpotrichoides*) and/or wheat glume blotch (*Septoria nodorum*). Preferred component (1) compounds for these compositions include 6-bromo-3-propyl-2-propyloxy-4(3*H*)-quinazolinone; 6,8-diiodo-3-propyl-2-propyloxy-4(3*H*)-quinazolinone; with

6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone being particularly preferred.

Preferred component (2) compounds for these compositions include 2,4-dihydro-5methoxy-2-methyl-4-[2-[[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, kresoxim-methyl, azoxystrobin, (E)-2-(methoxyimino)-N-methyl-2-(2-

phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and tebuconazole; with 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, fenpropimorph, flusilazole, epoxiconazole and propiconazole being particularly preferred. Preferably, the mole ratio of component (1) to component (2) for these 10

compositions is from about 4:1 to 1:4. Example compositions of this type include compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and 2,4dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one

in a mole ratio of the compound of Formula Ia to the compound of Formula IIa of from about 4:1 to 1:4, compositions comprising

15 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and fenpropimorph in a mole ratio of the compound of Formula Ia to fenpropimorph of from about 1:1 to 1:10, compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and flusilazole in a mole

ratio of the compound of Formula Ia to flusilazole of from about 15:1 to 1:15, compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and 20 tebuconazole in a mole ratio of the compound of Formula Ia to tebuconazole of from about 30:1 to 1:30, and compositions comprising

25

30

35

6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-

triazol-3-one; and flusilazole in a mole ratio of the compound of Formula Ia to the compound of Formula IIa plus flusilazole of from about 4:1 to 1:4 and the compound of Formula IIa to flusilazole has a mole ratio of from about 4:1 to 1:4.

This invention also provides a process for controlling at least one plant disease selected from wheat powdery mildew, wheat leaf rust, wheat foot rot, and wheat glume blotch which comprises applying to the plant or portion thereof to be protected, or to the plant seed or seedling to be protected, an effective amount of a fungicidal combination including component (1) and component (2) wherein the mole ratio of component (1) to component (2) is from about 30:1 to 1:30 (preferably from about 4:1 to 1:4). Component (1) can, for example, be applied at a rate of 0.2 g/ha or more. Typically component (1) is applied at a rate of 100 g/ha. Component (2) may be applied simultaneously (e.g., in the form of a composition comprising component (1) and component (2) in an appropriate mole ratio); or component (1) and component (2) can be applied separately in an appropriate mole ratio (e.g., as a tank mix).

Compositions wherein component (2) is selected from the group consisting of 2,4dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-

(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, kresoxim-methyl, azoxystrobin, (E)-2-(methoxyimino)-N-methyl-2-(2-

- 5 phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and tebuconazole; and wherein the mole ratio of component (1) to component (2) is from about 4:1 to 1:10 are considered especially useful for controlling wheat powdery mildew. Preferred component (1) compounds for these compositions include 6-bromo-3-propyl-2-propyloxy-4(3H)-quinazolinone;
- 10 6,8-diiodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; and 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; with 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone being particularly preferred. Preferred component (2) compounds for these compositions include 2,4-dihydro-5methoxy-2-methyl-4-[2-[[[[1-[3-
- 15 (trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, fenpropimorph, flusilazole, epoxiconazole and propiconazole. Example compositions of this type include compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and 2,4-dihydro-5-methoxy-2-
- methyl-4-[2-[[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-20 1,2,4-triazol-3-one in a mole ratio of the compound of Formula Ia to the compound of Formula IIa of from about 4:1 to 1:4, compositions comprising
 - 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and fenpropimorph in a mole ratio of the compound of Formula Ia to fenpropimorph of from about 1:1 to 1:10, compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and flusilazole in a mole
- 25 ratio of the compound of Formula Ia to flusilazole of from about 4:1 to 1:4, and compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; 2,4dihydro-5-methoxy-2-methyl-4-[2-[[[[1-[3-
 - (trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one; and flusilazole in a mole ratio of the compound of Formula Ia to the total of the
- 30 compound of Formula IIa plus flusilazole of from about 4:1 to 1:4 and a mole ratio of the compound of Formula IIa to flusilazole of from about 4:1 to 1:4.

This invention also provides a process for controlling wheat powdery mildew which comprises applying to the plant or portion thereof to be protected, or to the plant seed or seedling to be protected, an effective amount of a fungicidal combination including component (1) and component (2) wherein the mole ratio of component (1) to component (2) is from about 4:1 to 1:10 (preferably from about 4:1 to 1:4). Component (1) can, for example, be applied at a rate of 0.2 g/ha or more. Typically component (1) is applied at a rate of 100 g/ha. Component (2) may be applied

35

simultaneously (e.g., in the form of a composition comprising component (1) and component (2) in an appropriate mole ratio); or component (1) and component (2) can be applied separately in an appropriate mole ratio (e.g., as a tank mix).

Compositions wherein component (2) is selected from the group consisting of 2,4dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-5 (trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, kresoxim-methyl, azoxystrobin, (E)-2-(methoxyimino)-N-methyl-2-(2phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and tebuconazole; and wherein the mole ratio of component (1) to component (2) is from about 30:1 to 1:30 are considered especially useful for controlling wheat leaf rust. 10 Preferred component (1) compounds for these compositions include 6-bromo-3-propyl-2-propyloxy-4(3H)-quinazolinone; 6,8-diiodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; and 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; with 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone being particularly preferred being 15 particularly preferred. Preferred component (2) compounds for these compositions include 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, epoxiconazole, flusilazole and tebuconazole; with flusilazole and epoxiconazole being particularly preferred. Preferably, the mole ratio of component (1) to component (2) for 20 these compositions if from about 4:1 to 1:4. Example compositions of this type include compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and flusilazole in a mole ratio of the compound of Formula Ia to flusilazole of from about 15:1 to 1:15, compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and tebuconazole in a mole ratio of 25

This invention also provides a process for controlling wheat leaf rust which comprises applying to the plant or portion thereof to be protected, or to the plant seed or seedling to be protected, an effective amount of a fungicidal combination including component (1) and component (2) wherein the mole ratio of component (1) to component (2) is from about 30:1 to 1:30 (preferably from about 4:1 to 1:4). Component (2) can, for example, be applied at a rate of 12.5 g/ha or more. Typically component (2) is applied at a rate of 160 g/ha. Component (1) may be applied simultaneously (e.g., in the form of a composition comprising component (1) and component (2) in an appropriate mole ratio); or component (1) and component (2) can be applied separately in an appropriate mole ratio (e.g., as a tank mix).

the compound of Formula Ia to tebuconazole from about 30:1 to 1:30.

30

35

Compositions wherein component (2) is selected from the group consisting of 2,4dihydro-5-methoxy-2-methyl-4-[2-[[[[1-[3-

(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, kresoxim-methyl, azoxystrobin, (E)-2-(methoxyimino)-N-methyl-2-(2phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and tebuconazole; and wherein the mole ratio of component (1) to component (2) is from about 15:1 to 1:15 are considered especially useful for controlling wheat foot rot. Preferred component (1) compounds for these compositions include 6-bromo-3-propyl-2-propyloxy-4(3H)-quinazolinone; 6,8-diiodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; and

6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; with 10 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone being particularly preferred being particularly preferred. A particularly preferred component (2) compound for these compositions is flusilazole. Preferably, the mole ratio of component (1) to component (2) for these compositions if from about 4:1 to 1:4. Example compositions of this type include compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone 15 and flusilazole in a mole ratio of the compound of Formula Ia to flusilazole of from

about 15:1 to 1:15.

20

25

This invention also provides a process for controlling wheat foot rot which comprises applying to the plant or portion thereof to be protected, or to the plant seed or seedling to be protected, an effective amount of a fungicidal combination including component (1) and component (2) wherein the mole ratio of component (1) to component (2) is from about 15:1 to 1:15 (preferably from about 4:1 to 1:4). Component (1) can, for example, be applied at a rate of 5 g/ha or more. Typically component (1) is applied at a rate of 100 g/ha. Component (2) may be applied simultaneously (e.g., in the form of a composition comprising component (1) and component (2) in an appropriate mole ratio); or component (1) and component (2) can be applied separately in an appropriate mole ratio (e.g., as a tank mix).

Compositions wherein component (2) is selected from the group consisting of 2,4dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, 30 kresoxim-methyl, azoxystrobin, (E)-2-(methoxyimino)-N-methyl-2-(2phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and tebuconazole; and wherein the mole ratio of component (1) to component (2) is from about 30:1 to 1:30 are considered especially useful for controlling wheat glume blotch. Preferred component (1) compounds for these compositions include 35 6-bromo-3-propyl-2-propyloxy-4(3H)-quinazolinone; 6,8-diiodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; and 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone; with

6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone being particularly preferred being

10

15

20

25

30

particularly preferred. Preferred component (2) compounds for these compositions include flusilazole and tebuconazole; with flusilazole being particularly preferred. Preferably, the mole ratio of component (1) to component (2) for these compositions if from about 4:1 to 1:4. Example compositions of this type include compositions comprising 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and tebuconazole in a mole ratio of the compound of Formula Ia to tebuconazole from about 30:1 to 1:30

This invention also provide a process for controlling wheat glume blotch which comprises applying to the plant or portion thereof to be protected, or to the plant seed or seedling to be protected, an effective amount of a fungicidal combination including component (1) and component (2) wherein the mole ratio of component (1) to component (2) is from about 30:1 to 1:30 (preferably from about 4:1 to 1:4). Component (2) can, for example, be applied at 12.5 g/ha or more. Typically component (2) is applied at a rate of 160 g/ha. Component (1) may be applied simultaneously (e.g., in the form of a composition comprising component (1) and component (2) in an appropriate mole ratio); or component (1) and component (2) can be applied separately in an appropriate mole ratio, e.g., as a tank mix.

Synthesis of Compounds of Formula I

The compounds of Formula I where D is O can be prepared as described in International Patent Application WO 94/26722 and as shown in Scheme 1.

An anthranilic acid (2-aminobenzoic acid) of Formula 1 is condensed with an isothiocyanate of Formula R¹-NCS to form the 2-thioquinazolinedione of Formula 2. This condensation is preferably performed in the presence of a base such as triethylamine. S-Methylation of this compound affords the 2-methylthio-4(3H)-quinazolinone of Formula 3.

For the introduction of the R²O group, the 2-methylthio-4(3*H*)-quinazolinone of Formula 3 is treated with a mixture of a base, for example sodium hydride, in R²OH solvent. The reaction mixture is stirred at a temperature from about 0 °C to 120 °C for 1-120 hours. The desired 2-R²O-4(3H)-quinazolinone can be isolated from the reaction mixture by extraction into a water-immiscible solvent, and purified by chromatography or recrystallization. Similar synthetic procedures are described in U.S. 3,755,582, incorporated herein by reference.

Scheme 1

Anthranilic acids of Formula 1 are known or can be prepared by known methods. For example see, March, J. *Advanced Organic Chemistry*; 3rd ed., John Wiley:

New York, (1985), p 983. The isothiocyanates of Formula R¹-NCS can be prepared from the corresponding amine by treatment with thiophosgene as known in the art. For example, see *J. Heterocycl. Chem.*, (1990), 27, 407.

6-Iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone, Formula Ia, is a compound of Formula I where D is O, R^1 is propyl, R^2 is propyl, R^3 is iodine fixed in the 6 position and R^4 is hydrogen.

Synthesis of Compounds of Formula II

Compounds of Formula II where Q is Q-1 and X is OR¹¹ can be prepared as described in International Patent Application WO 95/14009 by treating a compound of Formula 4 with an appropriate alkyl transfer reagent in an inert solvent with or without additional acidic or basic reagents or other reagents (Scheme 2). Suitable solvents are selected from the group consisting of polar aprotic solvents such as acetonitrile, dimethylformamide or dimethylsulfoxide; ethers such as tetrahydrofuran, dimethoxyethane, or diethyl ether; ketones such as acetone or 2-butanone; hydrocarbons such as toluene or benzene; and halocarbons such as dichloromethane or chloroform.

10

15

5

Scheme 2

HO
$$A = N$$
 R^{12}
 R^{12}
 R^{12}
 R^{12}
 R^{12}

Method 1: $V-CH=N_2$ (V = H or $(CH_3)_3Si$)

Method 2: Cl₃C OR11; Lewis acid

Method 3: (R¹¹)₃O⁺ BF₄

Method 4: $(R^{11})_2SO_4$; $R^{11}OSO_2V$; or R^{11} -hal; optional base (hal = F, Cl, Br, or I) ($V = C_1 - C_6$ alkyl, $C_1 - C_6$ haloalkyl)

For example, compounds of Formula II where Q is Q-1 and X is OR^{11} can be prepared by the action of diazoalkane reagents of Formula 5 such as diazomethane (V = H) or trimethylsilyldiazomethane $(V = (CH_3)_3Si)$ on of Formula 4 (Method 1). Use of trimethylsilyldiazomethane requires a protic cosolvent such as methanol. For examples of these procedures, see *Chem. Pharm. Bull.*, (1984), 32, 3759.

As indicated in Method 2, compounds of Formula II where Q is Q-1 and X is OR¹¹ can also be prepared by contacting compounds of Formula 4 with alkyl

WO 98/33381 PCT/US98/01381

trichloroacetimidates of Formula 6 and a Lewis acid catalyst. Suitable Lewis acids include trimethylsilyl triflate and tetrafluoroboric acid. The alkyl trichloroacetimidates can be prepared from the appropriate alcohol and trichloroacetonitrile as described in the literature (J. Danklmaier and H. Hönig, *Synth. Commun.*, (1990), 20, 203).

Compounds of Formula II where Q is Q-1 and X is OR¹¹ can also be prepared from compounds of Formula 4 by treatment with a trialkyloxonium tetrafluoroborate (i.e., Meerwein's salt) of Formula 7 (Method 3). The use of trialkyloxonium salts as powerful alkylating agents is well known in the art (see U. Schöllkopf, U. Groth, C. Deng, Angew. Chem., Int. Ed. Engl., (1981), 20, 798).

5

10

15

20

25

Other alkylating agents which can convert compounds of Formula 4 to compounds of Formula II where Q is Q-1 and X is OR¹¹ are dialkyl sulfates such as dimethyl sulfate, haloalkyl sulfonates such as methyl trifluoromethanesulfonate, and alkyl halides such as iodomethane and propargyl bromide (Method 4). These alkylations can be conducted with or without additional base. Appropriate bases include alkali metal alkoxides such as potassium *tert*-butoxide, inorganic bases such as sodium hydride and potassium carbonate, or tertiary amines such as triethylamine, pyridine, 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU), and triethylenediamine. See R. E. Benson, T. L. Cairns, J. Am. Chem. Soc., (1948), 70, 2115 for alkylation examples using agents of this type.

Compounds of Formula II where Q is Q-1 and X is OR¹¹ can also be prepared by reaction of Formula 8 compounds with alkali metal alkoxides (R¹¹O-M⁺) (Scheme 3). The leaving group Lg¹ in the amides of Formula 8 are any group known in the art to undergo a displacement reaction of this type. Examples of suitable leaving groups include chlorine, bromine, and sulfonyl and sulfonate groups. Examples of suitable inert solvents are dimethylformamide or dimethylsulfoxide.

Scheme 3

$$\begin{array}{c} E & Y & Z \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & &$$

2.4-Dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, Formula IIa, is a compound of Formula II where Q is Q-1, E is 1,2-phenylene, A is N, G is N with the floating double bond attached to A, W is O, X is OR¹¹, R¹¹ is CH₃, R¹² is CH₃, Y is -CHR²⁵O-N=C(R¹⁷)-, R¹⁷ is CH₃, R²⁵ is H, Z is phenyl substituted with R¹⁹ and R¹⁹ is CF₃ fixed in the 3 position.

Azoxystrobin, Formula IIb, is a compound of Formula II where Q is Q-2, E is 1,2 phenylene, W¹ is O, X¹ is C, Y is -O-, Z is 4,6-pyrimidinyl substituted with R¹⁹, R¹⁹ is phenoxy substituted with R²¹ and R²¹ is cyano fixed in the 2 position.

Kresoxim-methyl, Formula IIc, is a compound of Formula II where Q is Q-2, E is 1,2-phenylene, W1 is O, X1 is N, Y is -CHR25O-, R25 is H, Z is phenyl substituted with R19 and R19 is CH3 fixed in the 2 position.

(E)-2-(methoxyimino)-N-methyl-2-(2-phenoxyphenyl)acetamide, Formula IId, is a compound of Formula II where Q is Q-2, E is 1,2-phenylene, W1 is NH, X1 is N, Y is -O-, Z is phenyl substituted with R¹⁹ wherein R¹⁹ is H.

Without further elaboration, it is believed that one skilled in the art using the preceding description can utilize the present invention to its fullest extent. The following Examples are, therefore, to be construed as merely illustrative.

Formulation/Utility 20

10

15

25

30

The fungicidal compositions of the present invention comprise an effective amount of a mixture of the compounds(s) of component (1) (e.g., 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone) and the compound(s) of component (2) (e.g., 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one, kresoxim-methyl, azoxystrobin, (E)-2-(methoxyimino)-N-methyl-2-(2phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and/or tebuconazole). The mixtures of this invention will typically be used as a formulation or composition with an agriculturally suitable carrier comprising at least

one of a liquid diluent, a solid diluent or a surfactant. The formulation or composition

. . 10

15

20

25

ingredients are selected to be consistent with the physical properties of the active ingredient, mode of application and environmental factors such as soil type, moisture and temperature. Useful formulations include liquids such as solutions (including emulsifiable concentrates), suspensions, emulsions (including microemulsions and/or suspoemulsions) and the like which optionally can be thickened into gels. Useful formulations further include solids such as dusts, powders, granules, pellets, tablets, films, and the like which can be water-dispersible ("wettable") or water-soluble. Active ingredient can be (micro)encapsulated and further formed into a suspension or solid formulation; alternatively the entire formulation of active ingredient can be encapsulated (or "overcoated"). Encapsulation can control or delay release of the active ingredient. Sprayable formulations can be extended in suitable media and used at spray volumes from about one to several hundred liters per hectare. High-strength compositions are primarily used as intermediates for further formulation.

The formulations will typically contain effective amounts of active ingredients, diluent and surfactant within the following approximate ranges which add up to 100 percent by weight.

Weight Percent **Active** Surfactant Ingredients Diluent 0-94 1-15 Water-Dispersible and Water-soluble 5-90 Granules, Tablets and Powders. 5-50 40-95 0-15 Suspensions, Emulsions, Solutions (including Emulsifiable Concentrates) 0-5 70--99 1-25 **Dusts** 0.01-99 0 - 155-99.99 **Granules and Pellets** 0-2 0-10 90-99 High Strength Compositions

Typical solid diluents are described in Watkins, et al., Handbook of Insecticide Dust Diluents and Carriers, 2nd Ed., Dorland Books, Caldwell, New Jersey. Typical liquid diluents are described in Marsden, Solvents Guide, 2nd Ed., Interscience, New York, 1950. McCutcheon's Detergents and Emulsifiers Annual, Allured Publ. Corp., Ridgewood, New Jersey, as well as Sisely and Wood, Encyclopedia of Surface Active Agents, Chemical Publ. Co., Inc., New York, 1964, list surfactants and recommended uses. All formulations can contain minor amounts of additives to reduce foam, caking, corrosion, microbiological growth and the like, or thickeners to increase viscosity.

Surfactants include, for example, polyethoxylated alcohols, polyethoxylated alkylphenols, polyethoxylated sorbitan fatty acid esters, dialkyl sulfosuccinates, alkyl sulfates, alkylbenzene sulfonates, organosilicones, *N*,*N*-dialkyltaurates, lignin

sulfonates, naphthalene sulfonate formaldehyde condensates, polycarboxylates, and polyoxyethylene/polyoxypropylene block copolymers. Solid diluents include, for example, clays such as bentonite, montmorillonite, attapulgite and kaolin, starch, sugar, silica, talc, diatomaceous earth, urea, calcium carbonate, sodium carbonate and bicarbonate, and sodium sulfate. Liquid diluents include, for example, water, *N*,*N*-dimethylformamide, dimethyl sulfoxide, *N*-alkylpyrrolidone, ethylene glycol, polypropylene glycol, paraffins, alkylbenzenes, alkylnaphthalenes, oils of olive, castor, linseed, tung, sesame, corn, peanut, cotton-seed, soybean, rape-seed and coconut, fatty acid esters, ketones such as cyclohexanone, 2-heptanone, isophorone and 4-hydroxy-4-methyl-2-pentanone, and alcohols such as methanol, cyclohexanol, decanol and tetrahydrofurfuryl alcohol.

10

15

20

25

35

Solutions, including emulsifiable concentrates, can be prepared by simply mixing the ingredients. Dusts and powders can be prepared by blending and, usually, grinding as in a hammer mill or fluid-energy mill. Suspensions are usually prepared by wet-milling; see, for example, U.S. 3,060,084. Granules and pellets can be prepared by spraying the active material upon preformed granular carriers or by agglomeration techniques. See Browning, "Agglomeration", *Chemical Engineering*, December 4, 1967, pp 147-48, *Perry's Chemical Engineer's Handbook*, 4th Ed., McGraw-Hill, New York, 1963, pages 8-57 and following, and International Patent Publication WO 91/13546. Pellets can be prepared as described in U.S. Patent No. 4,172,714. Water-dispersible and water-soluble granules can be prepared as taught in U.S. Patent No. 4,144,050, U.S. Patent No. 3,920,442 and German Patent Application DE 3,246,493. Tablets can be prepared as taught in U.S. Patent No. 5,180,587, U.S. Patent No. 5,232,701 and U.S. Patent No. 5,208,030. Films can be prepared as taught in Great Britain Patent Application GB 2,095,558 and U.S. Patent No. 3,299,566.

For further information regarding the art of formulation, see U.S. Patent No. 3,235,361, Col. 6, line 16 through Col. 7, line 19 and Examples 10-41; U.S. Patent No. 3,309,192, Col. 5, line 43 through Col. 7, line 62 and Examples 8, 12, 15, 39, 41, 52, 53, 58, 132, 138-140, 162-164, 166, 167 and 169-182;

30 U.S. Patent No. 2,891,855, Col. 3, line 66 through Col. 5, line 17 and Examples 1-4; Klingman, Weed Control as a Science, John Wiley and Sons, Inc., New York, 1961, pp 81-96; and Hance et al., Weed Control Handbook, 8th Ed., Blackwell Scientific Publications, Oxford, 1989.

In the following Examples, all percentages are by weight and all formulations are prepared in conventional ways.

Example A

Wettable Powder	
Active ingredient(s)	65.0%
dodecylphenol polyethylene glycol ether	2.0%
sodium ligninsulfonate	4.0%
sodium silicoaluminate	6.0%
montmorillonite (calcined)	23.0%.
Example B	
Granule	
Active ingredient(s)	10.0%
attapulgite granules (low volatile matter,	•
0.71/0.30 mm; U.S.S. No. 25-50 sieves)	90.0%.
Example C	
Extruded Pellet	
Active ingredient(s)	25.0%
anhydrous sodium sulfate	10.0%
crude calcium ligninsulfonate	5.0%
sodium alkylnaphthalenesulfonate	1.0%
calcium/magnesium bentonite	59.0%.
Example D	
Emulsifiable Concentrate	
Active ingredient(s)	20.0%
blend of oil soluble sulfonates	
and polyoxyethylene ethers	10.0%
	70.0%.
	Active ingredient(s) dodecylphenol polyethylene glycol ether sodium ligninsulfonate sodium silicoaluminate montmorillonite (calcined) Example B Granule Active ingredient(s) attapulgite granules (low volatile matter, 0.71/0.30 mm; U.S.S. No. 25–50 sieves) Example C Extruded Pellet Active ingredient(s) anhydrous sodium sulfate crude calcium ligninsulfonate sodium alkylnaphthalenesulfonate calcium/magnesium bentonite Example D Emulsifiable Concentrate Active ingredient(s) blend of oil soluble sulfonates

The compounds of this invention are useful as plant disease control agents. The present invention therefore further comprises a method for controlling plant diseases caused by fungal plant pathogens comprising applying to the plant or portion thereof to be protected, or to the plant seed or seedling to be protected, an effective amount of a compound of the invention or a fungicidal composition containing said compound. The compounds and compositions of this invention provide control of diseases caused by a broad spectrum of fungal plant pathogens in the Basidiomycete, Ascomycete, Oomycete and Deuteromycete classes. They are effective in controlling a broad spectrum of plant diseases, particularly foliar pathogens of ornamental, vegetable, field, cereal, and fruit crops. These pathogens include *Plasmopara viticola*, *Phytophthora infestans*, *Peronospora tabacina*, *Pseudoperonospora cubensis*, *Pythium aphanidermatum*, *Alternaria brassicae*, *Septoria nodorum*, *Septoria tritici*, *Cercosporidium personatum*, *Cercospora arachidicola*, *Pseudocercosporella herpotrichoides*, *Cercospora beticola*,

30

35

Botrytis cinerea, Monilinia fructicola, Pyricularia oryzae, Podosphaera leucotricha, Venturia inaequalis, Erysiphe graminis, Uncinula necatur, Puccinia recondita, Puccinia graminis, Hemileia vastatrix, Puccinia striiformis, Puccinia arachidis, Rhizoctonia solani, Sphaerotheca fuliginea, Fusarium oxysporum, Verticillium dahliae, Pythium aphanidermatum, Phytophthora megasperma, Sclerotinia sclerotiorum, Sclerotium rolfsii, Erysiphe polygoni, Pyrenophora teres, Gaeumannomyces graminis, Rynchosporium secalis, Fusarium roseum, Bremia lactucae and other generea and species closely related to these pathogens.

Mixtures of this invention can be further mixed with one or more other insecticides, fungicides, nematocides, bactericides, acaricides, growth regulators, 10 chemosterilants, semiochemicals, repellents, attractants, pheromones, feeding stimulants or other biologically active compounds to form a multi-component pesticide giving an even broader spectrum of agricultural protection. Examples of such agricultural protectants with which compounds of this invention can be formulated are: insecticides such as abamectin, acephate, azinphos-methyl, bifenthrin, buprofezin, carbofuran, 15 chlorfenapyr, chlorpyrifos, chlorpyrifos-methyl, cyfluthrin, beta-cyfluthrin, cyhalothrin, lambda-cyhalothrin, deltamethrin, diafenthiuron, diazinon, diflubenzuron, dimethoate, esfenvalerate, fenoxycarb, fenpropathrin, fenvalerate, fipronil, flucythrinate, tau-fluvalinate, fonophos, imidacloprid, isofenphos, malathion, metaldehyde, methamidophos, methidathion, methomyl, methoprene, methoxychlor, methyl 7-chloro-20 2,5-dihydro-2-[[N-(methoxycarbonyl)-N-[4-(trifluoromethoxy)phenyl]amino]carbonyl]indeno[1,2-e][1,3,4]oxadiazine-4a(3H)carboxylate (DPX-JW062), monocrotophos, oxamyl, parathion, parathion-methyl, permethrin, phorate, phosalone, phosmet, phosphamidon, pirimicarb, profenofos, rotenone, sulprofos, tebufenozide, tefluthrin, terbufos, tetrachlorvinphos, thiodicarb, 25 tralomethrin, trichlorfon and triflumuron; fungicides such as benomyl, blasticidin-S, Bordeaux mixture (tribasic copper sulfate), captafol, captan, carbendazim, chloroneb, chlorothalonil, copper oxychloride, copper salts, cymoxanil, cyprodinil (CGA 219417), diclomezine, dicloran, dimethomorph, dodine, edifenphos, famoxadone, fenpiclonil, fluazinam, flutolanil, folpet, fosetyl-aluminum, furalaxyl, iprobenfos, iprodione, 30 isoprothiolane, kasugamycin, mancozeb, maneb, mepronil, metalaxyl, S-methyl 7-benzothiazolecarbothioate (CGA 245704), myclobutanil, neo-asozin (ferric methanearsonate), oxadixyl, pencycuron, probenazole, prochloraz, pyrifenox, pyroquilon, quinoxyfen, spiroxamine (KWG4168), sulfur, thiabendazole, thiophanate-methyl, thiram, tricyclazole, validamycin and vinclozolin; nematocides 35 such as aldoxycarb and fenamiphos; bactericides such as streptomycin; acaricides such as amitraz, chinomethionat, chlorobenzilate, cyhexatin, dicofol, dienochlor, etoxazole, fenazaquin, fenbutatin oxide, fenpropathrin, fenpyroximate, hexythiazox, propargite,

pyridaben and tebufenpyrad; and biological agents such as *Bacillus thuringiensis*, *Bacillus thuringiensis* delta endotoxin, baculovirus, and entomopathogenic bacteria, virus and fungi.

In certain instances, combinations with other fungicides having a similar spectrum of control but a different mode of action will be particularly advantageous for resistance management.

5

10

15

20

25

30

35

Plant disease control is ordinarily accomplished by applying an effective amount of the composition of this invention either pre— or post—infection, to the portion of the plant to be protected such as the roots, stems, foliage, fruit, seeds, tubers or bulbs, or to the media (soil or sand) in which the plants to be protected are growing. The composition can also be applied to the seed to protect the seed and seedling.

Rates of application for this composition can be influenced by many factors of the environment and should be determined under actual use conditions. Foliage can normally be protected when treated at a rate of from less than 1 g/ha to 5,000 g/ha of aggregate active ingredient. Aggregate active ingredient is defined as the total combined weight of active ingredients. Seed and seedlings can normally be protected when seed is treated at a rate of from 0.1 to 10 g of aggregate active ingredient per kilogram of seed.

The following Examples demonstrate the composition and method of the present invention and provide experimental evidence for synergy between the compound of Formula Ia and flusilazole in preventative control of wheat leaf rust caused by *Puccinia recondita*, wheat foot rot caused by *Pseudocercosporella herpotrichoides*, and wheat glume blotch caused by *Septoria nodorum*. The experimental also provides evidence for synergy between the compound of Formula Ia and tebuconazole in preventative control of wheat leaf rust and wheat glume blotch. The experimental also provides evidence for synergy between the compound of Formula Ia and flusilazole in curative control of wheat powdery mildew caused by *Erysiphe graminis f. sp. tritici* and wheat leaf rust.

The experimental also provides evidence for synergy between the compounds of Formula Ia, Formula IIa and flusilazole in preventative control of wheat foot rot. The experimental also provides evidence for synergy between the compounds of Formula Ia, IIa and flusilazole in curative control of wheat powdery mildew. The pathogen control protection afforded by these compositions is not limited, however, to these species.

The following TESTS demonstrate the control efficacy of compounds of this invention on specific pathogens. The pathogen control protection afforded by the compounds is not limited, however, to these species.

BIOLOGICAL EXAMPLES OF THE INVENTION

Test compounds were first dissolved in acetone in an amount equal to 50% of the final volume and then suspended at a concentrations from 0.08 to 200 ppm in purified water containing 250 ppm of the surfactant Trem® 014 (polyhydric alcohol esters). The resulting test suspensions were then used in the following test protocols. Spraying these test suspensions to the point of run-off on the test plants is the equivalent of a rate of 500 g/ha.

5

10

20

25

30

35

Protocol #1 - WPM Preventive

The test compounds were sprayed to the point of run-off on wheat seedlings. The following day the seedlings were inoculated with a spore dust of Erysiphe graminis f. sp. tritici, (the causal agent of wheat powdery mildew). Seedlings were incubated in a growth chamber at 20 °C for 6 days, after which disease ratings were made.

Protocol #2 - WPM Curative

Wheat seedlings were inoculated with a spore dust of Erysiphe graminis f. sp. tritici, (the causal agent of wheat powdery mildew). The following day a test 15 compounds were sprayed to the point of run-off on and seedlings incubated in a growthchamber at 20 °C for 7 days, after which disease ratings were made.

Protocol #3 WLR Preventive

The test compounds were sprayed to the point of run-off on wheat seedlings. The following day the seedlings were inoculated with a spore suspension of Puccinia recondita (the causal agent of wheat leaf rust) and incubated in a saturated atmosphere at 20 °C for 24 h, and then moved to a growth chamber at 20 °C for 6 days, after which disease ratings were made.

Protocol #4 - WLR Curative

Wheat seedlings were inoculated with a spore suspension of Puccinia recondita (the causal agent of wheat leaf rust). The following day the test compounds were sprayed to the point of run-off on and incubated in a saturated atmosphere at 20 °C for 24 h, then moved to a growth chamber at 20 °C for 6 days, after which disease ratings were made.

Protocol #5 - WFR Preventive

The test compounds were sprayed to the point of run-off on wheat seedlings. The following day the seedlings were inoculated with a spore suspension of Pseudocercosporella herpotrichoides (the causal agent of wheat eye spot or wheat foot rot) and incubated in a saturated atmosphere at 20 °C for 72 h, and then moved to a growth chamber at 20 °C for 6 days, after which disease ratings were made.

Protocol #6 - WGB Preventive

The test compounds were sprayed to the point of run-off on wheat seedlings. The following day the seedlings were inoculated with a spore suspension of Septoria

10

15

20

nodorum (the causal agent of wheat glume blotch) and incubated in a saturated atmosphere at 20 °C for 48 h, and then moved to a growth chamber at 20 °C for 6 days, after which disease ratings were made.

Protocol #7 - WGB Curative

Wheat seedlings were inoculated with a spore suspension Septoria nodorum (the causal agent of wheat glume blotch). Two days later test compounds were sprayed to the point of run-off on and seedlings incubated in a saturated atmosphere at 20 °C for 48 h, then moved to a growth chamber at 20 °C for 6 days, after which disease ratings were made.

Protocol #8 - WPM Preventative

The test suspension was sprayed to the point of run-off on wheat seedlings. The following day the seedlings were inoculated with a spore dust of *Erysiphe graminis* f. sp. tritici, (the causal agent of wheat powdery mildew) and incubated in a growth chamber at 20 °C for 7 days, after which disease ratings were made.

Results for protocols 1 - 8 are given in Tables A-G. In the tables, a rating of 100 indicates 100% disease control and a rating of 0 indicates no disease control (relative to the controls). An (nt) indicates no test results and (--) indicates no estimate of activity. In Tables A-G rates are given in parts per million (ppm) and/or grams active ingredient per hectare (g a.i./ha). The term "Actual" stands for the experimental value. The abbreviation "Exp." stands for "Expected" (i.e., the predicted activity, p, from the Colby equation).

TABLEA

Synergistic Effects of Compound Ia in Combination with Flusilazole, Tebuconazole and Prochloraz for Preventive Control of Wheat Powdery Mildew (WPM), Wheat Leaf Rust (WLR), Wheat Foot Roi (WFR) and Wheat Glume Blotch (WGB)

	ଚ ।	Exp.	ł	1.	1	:	ŀ	:	1	1	I	i	0	0	0	09	83	100
	$\overline{\text{WGB}^{(4)}}$	Actual	0	0	o	0	0	0	0	09	83	100	0	0	0	43	91	66
 1		<u>,</u>	1	1	ł	ł	1	ł	1	1	ŀ	1	Ξ	25	4.	2(10	10
ase Contro	WFR	Actual	0	0	0	0	16	54	45	20	100	100	0	∞	100	81	100	100
Percent Disease ((2)	Exp.	:	i	:		ŀ	ì	ł	ŀ	ı	ŀ	18	17	88	27	79	100
Pel	WLF	Actual Exp.	0	6	0.	0	18	6	88	27	79	100	6	18	88	32	86	66
	(L)	9	5															
	WPM ⁽¹⁾	Actual ⁽⁵⁾	100	100	100	100	06	66	100	62	35	81	100	100	100	100	nt(8)	nt
		Ratio											1.18:1	1.18:1	1.18:1	1:1.69	2.95:1	11.81:1
	Rates	(g a.i./ha)	0.2	-	5	25.	0.2	-	5	2.5	12.5	50	0.2 + 0.2	1+1	5+5	2.5 + 5	12.5 + 5	50 + 5
	Rates	(mdd)	80.0	0.4	7	10	80.0	0.4	2		· •	20	0.08 + 0.08	0.4 + 0.4	2+2	1+2	5+2	20+2
		Cmpd 2						-						Ia				Ia
	Test	Cmpd 1	Ia	Ia	Ia	la	Flusilazole											

	3(4)	Exp.	09	83	100	1	1	i	1	:	55	96	100	;	55	96	100	:	:	:	!	ł
	WG	Actual	38	94	100	nt	55	96	100		29	100	100	nt	84	100	100	nt	Ħ	nt	Ħ	Ħ
71	<u></u>	Exp.	-05	100	100	ł	ł	1	1	1	ł	ł	1	ł	ł	ł	ł	ŀ	ŀ	1	37	100
Percent Disease Control	WFR ⁽³⁾	Actual	96	100	100	nt	nt	nt	Ħ	Ħ	nt	n	nt	nt	nţ	nt	nt	37	100	100	37	100
rcent Dise	R ⁽²⁾	Exp.	27	79	100	ł	ı	i	I	32	100	100	ł	32	100	100	ŀ	i	ŀ	1	i	ı
ଥ	WLR ⁽²⁾	Actual	70	100	100	32	100	100	nt	86	100	100		9/	66	100	Ħ	nt	nt	nţ	nt	nt
	\	Exp. (6)	ł	ŀ	1	ì	ŀ	1	-}	ŀ	1	1	ì	ŀ	ł	ł	ł	1		}	ŀ	;
•	$WPM^{(1)}$	Actual ⁽⁵⁾	nt	n	Ħ	ti	Ħ	ti	nt	nt	nt	Ħ	nt	ti	ta	tu .	· nt	nt	nt	nt	пt	, nt
	Mole	Ratio	1:8.47	1:1.69	2.36:1					1:4.13	1:1.21	6.06:1	24.23:1	1:20.63	1:4.13	1:1.21	4.85:1				1:5	1:1
	Rates	(g a.i./ha)	2.5 + 25	12.5 + 25	50 + 25	-	. 5	25	100	1+5	5+5	25 + 5	100 + 5	1 + 25	5 + 25	25 + 25	100 + 25	-	کی	25	1+5	5+5
	Rates	(mdd)	1 + 10	5 + 10	20 + 10	0.4	2	10	40	0.4 + 2	2+2	10 + 2	40 + 2	0.4 + 10	2 + 10	10 + 10	40 + 10	0.4	7	10	0.4 + 2	2+2
	Test	Cmpd 2	Ia	Ia	Ia	ł	1	٠ ،	i	Ia	Ia	Ia	Ia	ľa	Ia	Ia	Ia	ł	ŀ	i	Ia	Ia
	Test	Cmpd 1	Flusilazole	Flusilazole	Flusilazole	Cebuconazole	rebuconazole	rebuconazole	[ebuconazole	[ebuconazole	[ebaconazole	Pebuconazole	Febuconazole	Febuconazole	Febuconazole	Febuconazole	Tebuconazole	Prochloraz	Prochloraz	Prochloraz	Prochloraz	Prochloraz

	(£)	Exp.	ł	ł	1	1	
	$\overline{\text{WGB}^{(4)}}$						
ᅙ	R ₍₃₎	Exp.	100	37	100	100	
ease Contr	$\overline{\text{WLR}^{(2)}}$	Actual	100	37	1:00	100	
rcent Dis	(Z)	Exp.	:	ŀ	1	ı	
Pe	WE	Actual	nţ	nt	nt	Ħ	
	Œ.	Exp. (6)	1	:	ŀ	1	
	WPI	Actual ⁽⁵⁾ Exp. ⁽⁶⁾	Ħ	n	nt	nt	
	Mole	Ratio	5:1	1:25	1:5	1:1	
	Rates	(g a.i./ha)	25 + 5	1+25	5 + 25	25 + 25	
	Rates	(mdd)	10+2	0.4 + 10	2 + 10	10 + 10	
	Test	Cmpd 2	. Ia	Ia	Ia	Ia	
	Test	Cmpd 1	Prochloraz	Prochloraz	Prochloraz	Prochloraz	

(1) Wheat powdery mildew activity - Protocol #1

(2) Wheat leaf rust activity - Protocol #3

(3) Wheat foot rot activity - Protocol # 5

(4) Wheat glume blotch activity - Protocol # 6

(5) Actual activity observed in the test

(6) Predicted activity

(7) No estimate - (--)

(8) Not tested (nt)

WPM), Syn

ldew (W			Exp.	!		!	!	!	:	:		1	•	1	!	ł	15	23	72
y. Mi		$WGB^{(3)}$	邱	•	•	•		•										,	
neat Powder	1	MG	Actual	0	0	10	0	0	0	S	0	15	69	pt	nt	nt	0	43	71
ntrol of Wl	ase Contro	R ⁽²⁾	Exp.	ŀ	1	ı	ł	ł	1	ŀ	ł	;	:	:	1	l	43	84	100
Curative Co	Percent Disease Control	WLR ⁽²⁾	Actual	0	0	0	0	0	0	43	80	84	100	ㅂ	Ħ	nt	15	100	66
nazole for (Glume Blot	ᆈ	() [Exp.	1	:	ŀ	1	1,	ı	1	:	i	ł	89	82	100	ł	:	;
or Tebuco		WPM ⁽¹⁾	Actual	0	99	09	nţ	89	73	89	100	91	100	86	100	100	nt	ıţ	nt
f h Flusilazolo tust (WLR)		Mole	Ratio											1.18:1	1.18:1	1.18:1	1:1.69	2.95:1	11.81:1
in Combination with Flusilazole or Tebuconazole for Curative Control of Wheat Powdery Mildew (W Wheat Leaf Rust (WLR) and Wheat Glume Blotch (WGB)		Rates	(g a.i./ha)	0.2	_	. 2	25	0.2		2.5	\$	12.5	20	0.2 + 0.2	1+1	5+5	2.5 + 5	12.5 + 5	50 + 5
ound Ia in Co		Rates	(mdd)	0.08	0.4	7	10	0.08	0.4	1	7	5	20	0.08 + 0.08	0.4 + 0.4	2+2	1+2	5+2	20+2
ects of Comp		Test	Cmpd 2	.	٠,١	i	;	1	:	1	ł	ŀ	ŀ	Ia	Ia	Ia	Ia	Ia	Ia
Synergistic Effects of Compound Ia		Test	Cmpd 1	Ia	Ia	Ia	Ia	Flusilazole											

		Exp.	0	15	69	ŀ	;	ł	ŀ	:	49	55	91	:	43	20	90
	WGB(3)								•								
힑		Actu	0	29	81	Ħ	43	. 20)6 ,	nt	29	71	96	n)[7	10
ase Contr	R ⁽²⁾	Exp.	43	84	100	ì	1	ŀ	1	40	100	100	100	40	100	100	100
Percent Disease Contro	WLR ⁽²⁾	<u>Actual</u>	71	100	100	40	100	100	100	100	100	100	100	100	100	100	100
ш,	Œ.	Exp.	•	ŀ	ŀ	1	ì	ł	ł	ł	ı	1	ł	1	ŀ	:	
	WPM ⁽¹⁾	Actual	nt	nt	nţ	, ta	nt	nt	nt	nt	Ħ	Ħ	nt	tt	Ħ	Ħ	Ħ
	Mole	Ratio	1:8.47	1:1.69	2.36:1					1:4.13	1:1.21	6.06:1	24.23:1	1:20.63	1:4.13	1:1.21	4.85:1
	Rates	(g a.i./ha)	2.5 +25	12.5 + 25	50 + 25	1	5	25	100	1+5	5+5	25 + 5	100 + 5	1 + 25	5+25	25 + 25	100 + 25
	Rates	(mdd)	1 + 10	5 + 10	20 + 10	0.4	7	10	40	0.4+2	2+2	10+2	40+2	0.4 + 10	2 + 10	10 + 10	40 + 10
	Test	Cmpd 2	Ia	Ia	Ia	ŀ	1	٠ ١	ł	Ia	_						
	Test	Cmpd 1	Flusilazole	Flusilazole	Flusilazole	Tebuconazole	Tehnconszole										

Wheat powdery mildew - Protocol # 2
 Wheat leaf rust - Protocol # 4
 Wheat glume blotch - Protocol # 7

Synergistic Effects of Compound Ia in Combination with Compound IIa for Preventive Control of Wheat Powdery Mildew (WPM), Wheat Leaf Rust (WLR), Wheat Foot Rot (WFR) and Wheat Glume Blotch (WGB) TABLEC

B(4)	Exp.	ł	ŀ	1	1	}	1	0	0	0
WG	Actual Exp.	0	0	0	0	0	0	0	0	0
R(3)	Exp.	ŀ	ł	i	l	i	ł	∞	76	% %
WE	Actual Exp. Actual Exp. A	0	o .	0	∞	26	38	0	∞	61
R ⁽²⁾	Exp.	:	:	;	:	. !	ŀ	54	92	91
M	Actual	0	6	0	54	91	66	54	93	66
() W	Exp.	ł	ŀ	i	ŀ	ŀ	ł	100	100	100
WP	Actual Exp. A	100	100	100	09	88	100	66	100	100
	Ratio								1:1.13	
Rates	(g a.i./ha)	0.2	1	. 8	0.2	1	ν.	0.2 + 0.2	1 + 1	5+5
Rates	(mdd)	0.08	0.4	2	0.08	0.4	2	0.08 + 0.08	0.4 + 0.4	2+2
	Cmpd 3									
Test	Cmpd 2	1	!		;	:	i	Ia	Ia	Ia
Test	Cmpd 1	la	Ia	Ia	IIa	IIa	IIa	IIa	IIa	IIa

Wheat powdery mildew activity - Protocol #1
 Wheat leaf rust activity - Protocol # 3

(4) Wheat glume blotch activity - Protocol # 6

(3) Wheat foot rot activity - Protocol # 5

TABLED

Synergistic Effects of Compound Ia in Combination with Compound IIa on Curative Control of Wheat Powdery Mildew (WPM), Wheat Leaf Rust (WLR) and Wheat Glume Blotch (WGB)

	B ⁽³⁾	Exp.	1	1	;	ŀ	ı	1	13	0	22
힑	$\overline{\text{WGB}^{(3)}}$	Actual	0	0	10	13	0	13	0	13	53
se Contr	_ල ූ	Exp.	1	ł	ŀ	ł	ł	ł	0	0	0
ercent Disease				- 0							
집	() ()	Exp.	1	ì	1.	ì	1	1	0	88	100
	WP	Actual	0	95	09	0	73	100	21	66	100
	Mole	Ratio							1:1.13	1:1.13	1:1.13
	Rates	(g a.i./ha)	0.2	1	5	0.2	1	ν,	0.2 + 0.2	1+1	5+5
	Rates	(mdd)	0.08	0.4	2	0.08	0.4	7	0.08 + 0.08	0.4 + 0.4	2+2
	Test	Cmpd 3	1	ŀ	!	ł	ł	1	:	ı	:
	Test	Cmpd 2	!	- 1		ł	ŀ	;	Ia	Ia	Ia
	Test	Cmpd 1	Ia	Ia	Ia	IIa	IIa	IIa	IIa	IIa	IIa

(1) Wheat powdery mildew - Protocol # 2

⁽²⁾ Wheat leaf rust - Protocol # 4

⁽³⁾ Wheat glume blotch - Protocol # 7

TABLEE

Synergistic Effects of Compound Ia / Compound IIa / Flusilazole combinations for Preventive Control of Wheat Powdery Mildew (WPM), Wheat Leaf Rust (WLR), Wheat Foot Rot (WFR) and Wheat Glume Blotch (WGB)

							44	•						
	B(4)	Exp.	ł	ł	1	ŀ	ŀ	ŀ	ł	1	1	0	0	0
	WGB ⁽⁴⁾	Actual	0	0	0	0	0	0	0	0	0	0	o [.]	47
뎨	ଞ୍ଚା	Exp.	ŀ	;					ŀ		ł	16	24	
ease Cont	WFR ⁽³⁾	Actual	0	0	0	∞	. 26	38	16	54	45	∞	62	100
Percent Disease	_ව ූ	Exp.	1	:	:			+			1		100	100
Perc	WLR ⁽²⁾	Actual	0	6	0	54	. 91	66	18	6	88	28	93	66
	ĵ.	Exp.	1	. }	ł	;	ł	1	ŀ	ł	ł	100	100	100
	WPM ⁽¹⁾	Actual	100	100	100	09	88	100	06	66	100	100	100	100
	Mole	Ratio										1.33:1.13:1	1.33:1.13:1	1.33:1.13:1
	Rates	(g a.i./ha)	0.2	1	'	0.2	1	ری	0.2	• <u>-</u> -		0.2 + 0.2 + 0.2	1 + 1 + 1	5+5+5
	Rates	(mdd)	0.08	0.4	2		-			•		0.08	0.4	2+2+2
		•		;										
	Test	Cmpd 2	:	1	;	ł		ţ	ł	i	ŀ	Ia	Ia	Ia
	Test	Cmpd 1	Ia	Ia	Ia	IIa	IIa	IIa	Plusilazole	Flusilazole	Flusilazole	Flusilazole	Flusilazole	Flusilazole

Wheat powdery mildew activity - Protocol #1
 Wheat leaf rust activity - Protocol # 3

(4) Wheat glume blotch activity - Protocol # 6

(3) Wheat foot rot activity - Protocol # 5

Synergistic Effects of Compound Ia / Compound IIa / Flusilazole on Curative Control of Wheat Powdery Mildew (WPM), Wheat Leaf Rust (WLR) and Wheat Glume Blotch (WGB) TABLEF

							-							
	WGB ⁽³⁾	Exp.	1	ł	1	ł	ł	ŀ	;	ł	1	13	0	22
	WG	Actual Exp	0	0	10	13	0	13	0	0	0	40	0	13
e Contro	(g)	Exp.	ł	i	i	ł	1	ì	:	!	ŀ	0	0	80
Percent Disease Control	WLR ⁽²⁾	<u>Actual</u>	0	0	0	0	0	0	0	0	80	0	0	98
Perc	Ę.	Exp.	1	ŀ	ŀ	ł	ł		ł		1	89	100	100
	WPM ⁽¹⁾	Actual	0	99	09	0	73	100	89	73	100	96	100	100
	Mole	Ratio										1.33:1.13:1	1.33:1.13:Ì	1.33:1.13:1
	Rates	(g a.i./ha)	0.2	, -	5	0.2	1	ς,	0.2		ς.	0.2 + 0.2 + 0.2	1 + 1 + 1	5+5+5
	Rates	(mdd)	0.08	0.4	2	0.08	0.4	2	0.08	0.4	2	0.08 + 0.08 + 0.08	0.4 + 0.4 + 0.4	2+2+2
	Test	Cmpd 3		ŀ	ł	ŧ	ł	ł	1	l	ł	IIa	IIa	IIa
	Test	Cmpd 2	:	-1	ŀ	ı	1	1	1		ł	Ia	• Ia	Ia
	Test	Cmpd 1	Ia	Ia	Ia	IIa	IIa	IIa	Flusilazole	Flusilazole	Flusilazole	Flusilazole	Flusilazole	Flusilazole

(1) Wheat powdery mildew - Protocol #2

⁽²⁾ Wheat leaf rust - Protocol #4

⁽³⁾ Wheat glume blotch - Protocol #7

Synergistic Effects of Compound Ia in Combination with Fenpropimorph for Preventative Control of Wheat Powdery Mildew (WPM)(1)

	$\overline{\text{Exp.}(3)}$		(4)	:	ŀ	ŀ	١.	ŀ	100	9.7.6	72				st		
	Mean		100	96	72	. 8/	41	0	100	66	87				n the te		
PM(2)	Test 3		100	26	34	80	13	Ο,	100	86	82	52			served		
ol of W	Test 1 Test 2 Test 3		100	95	87	9/	9/	nt(5)	100	100	Ħ	31			ivity ob	te - ()	
% Control of WPM(2)	Test 1		100	96	94	79	34	0	100	66	92	19			(2) Actual activity observed in the test	(4) No estimate - ()	
	Mole	Ratio							6.1:1	4.9:1	6.1:1			,	(2) A	(4) N	
	Rates (g a.i./ha)		20	Ś	1	100	20	S	100 + 20	20+5	5 + 1				/ - Protocol #8		,
	Cmpd 2		1	ŀ	ŀ	ı	ŀ	ł	Ia	Ia	Ia		٠	•	nildew activity		
	Cmpd 1		Ia	Ia	. Ia	Fenpropimorph	Fenpropimorph	Fenpropimorph	Fenpropimorph	Fenpropimorph	Fenpropimorph	Untreated	(% area diseased	Foliage)	(1) Wheat powdery mildew activity - Protocol #8	(3) Predicted activity	(5) Not tested (nt)

CLAIMS

What is claimed is:

- A fungicidal composition comprising:
- (1) at least one compound selected from the quinazolinones of Formula I,
- N-oxides, and agriculturally suitable salts thereof,

$$R^3$$
 R^1
 R^4
 R^4
 R^2

wherein

D is O or S;

R¹ is C₁-C₁₀ alkyl; C₃-C₅ cycloalkyl; C₃-C₁₀ alkenyl; C₃-C₁₀ alkynyl; 10 C₁-C₁₀ haloalkyl; C₃-C₁₀ haloalkenyl; C₃-C₁₀ haloalkynyl; C_2 - C_{10} alkoxyalkyl; C_2 - C_{10} alkylthioalkyl; C_2 - C_{10} alkylsulfinylalkyl; C_2 - C_{10} alkylsulfonylalkyl; C_4 - C_{10} cycloalkylalkyl; C₄-C₁₀ alkenyloxyalkyl; C₄-C₁₀ alkynyloxyalkyl; C₄-C₁₀ (cycloalkyl)oxyalkyl; C₄-C₁₀ alkenylthioalkyl; C₄-C₁₀ alkynylthioalkyl; 15 C₄-C₁₀ (cycloalkyl)thioalkyl; C₂-C₁₀ haloalkoxyalkyl; C₄-C₁₀ haloalkenyloxyalkyl; C₄-C₁₀ haloalkynyloxyalkyl; C₄-C₁₀ alkoxyalkenyl; C₄-C₁₀ alkoxyalkynyl; C₄-C₁₀ alkylthioalkenyl; C₄-C₁₀ alkylthioalkynyl; C₄-C₁₀ trialkylsilylalkyl; C₁-C₁₀ alkyl substituted with NR5R6, nitro, cyano or phenyl optionally substituted with R8, R9 and 20 R^{10} ; C_1 - C_{10} alkoxy; C_1 - C_{10} haloalkoxy; C_1 - C_{10} alkylthio; C1-C10 haloalkylthio; or pyridinyl, furanyl, thienyl, naphthalenyl, benzofuranyl, benzothienyl or quinolinyl each optionally substituted with R⁸, R⁹ and R¹⁰;

R² is C₁-C₁₀ alkyl; C₃-C₇ cycloalkyl; C₃-C₁₀ alkenyl; C₃-C₁₀ alkynyl; 25 C₁-C₁₀ haloalkyl; C₃-C₁₀ haloalkenyl; C₃-C₁₀ haloalkynyl; C₂-C₁₀ alkoxyalkyl; C₂-C₁₀ alkylthioalkyl; C₂-C₁₀ alkylsulfinylalkyl; C₂-C₁₀ alkylsulfonylalkyl; C₄-C₁₀ cycloalkylalkyl; C₄-C₁₀ alkenyloxyalkyl; C₄-C₁₀ alkynyloxyalkyl; C₄-C₁₀ (cycloalkyl)oxyalkyl; C₄-C₁₀ alkenylthioalkyl; C₄-C₁₀ alkynylthioalkyl; 30 C₄-C₁₀ (cycloalkyl)thioalkyl; C₂-C₁₀ haloalkoxyalkyl; C₄-C₁₀ haloalkenyloxyalkyl; C₄-C₁₀ haloalkynyloxyalkyl; C₄-C₁₀ alkoxyalkenyl; C₄-C₁₀ alkoxyalkynyl; C₄-C₁₀ alkylthioalkenyl;

20

30

- R^1 and R^2 are taken together to form - $CH_2(CH_2)_qCH_2$ -; q is 0, 1, 2, 3 or 4;
 - R³ is halogen, C₁-C₈ alkyl, C₃-C₈ cycloalkyl, C₂-C₈ alkenyl, C₂-C₈ alkynyl, C₁-C₈ haloalkyl, C₃-C₈ haloalkenyl, C₃-C₈ haloalkynyl, C₁-C₈ alkoxy, C₁-C₈ haloalkoxy, C₃-C₈ alkenyloxy, C₃-C₈ alkynyloxy, C₁-C₈ alkylthio, C₃-C₈ alkenylthio, C₁-C₈ alkylsulfinyl, C₁-C₈ alkylsulfinyl, C₂-C₈ alkylsulfinylalkyl, C₂-C₈ alkylsulfonylalkyl, C₄-C₈ cycloalkylalkyl, C₃-C₈ trialkylsilyl, NR⁵R⁶, C₅-C₈ trialkylsilylalkynyl or phenyl optionally substituted with at least one R⁷;
- 15 R^4 is hydrogen, halogen, C_1 - C_4 alkyl, C_1 - C_4 haloalkyl, C_1 - C_4 alkoxy or C_1 - C_4 haloalkoxy;
 - each R⁵ is independently hydrogen, C₁-C₄ alkyl or phenyl optionally substituted with at least one R⁷;
 - each R⁶ is independently hydrogen, C₁-C₈ alkyl or phenyl optionally substituted with at least one R⁷; or
 - when R⁵ and R⁶ are attached to the same nitrogen atom, R⁵ and R⁶ can be taken together to form -CH₂CH₂CH₂CH₂-, -CH₂(CH₂)₃CH₂-, -CH₂CH₂OCH₂CH₂-, -CH₂CH(Me)CH₂- or -CH₂CH(Me)OCH(Me)CH₂-;
- each R⁷ is independently halogen, C₁-C₄ alkyl, C₁-C₄ alkoxy, C₁-C₄ haloalkyl, nitro or cyano;
 - R⁸ is C₁-C₆ alkyl, C₁-C₆ alkoxy, C₁-C₆ haloalkyl, halogen, C₂-C₈ alkynyl, C₁-C₆ alkylthio, phenyl or phenoxy each optionally substituted with at least one R⁷, cyano, nitro, C₁-C₆ haloalkoxy, C₁-C₆ haloalkylthio, C₂-C₆ alkenyl, C₂-C₆ haloalkenyl, acetyl, CO₂Me or N(C₁-C₂ alkyl)₂;
 - each R⁹ is independently methyl, ethyl, methoxy, methylthio, halogen or trifluoromethyl; and
 - each R10 is independently halogen;
- (2) at least one compound selected from (a) compounds acting at the bc₁ complex
 of the fungal mitochondrial respiratory electron transfer site and (b) compounds that control fungal disease by inhibiting sterol biosynthesis; and optionally

PCT/US98/01381 WO 98/33381

49

(3) at least one of a surfactant, a solid diluent or a liquid diluent; wherein component (1) and component (2) are present in a fungicidally effective amount and the mole ratio of component (1) to component (2) is from about 30:1 to 1:30.

The fungicidal composition of Claim 1 comprising (1) at least one compound selected from the group consisting of

6-bromo-3-propyl-2-propyloxy-4(3H)-quinazolinone,

- 6,8-diiodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and
- 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone;

5

10

15

20

25

30

- and (2) at least one compound selected from the group consisting of
- 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4triazol-3-one, kresoxim-methyl, azoxystrobin, (E)-2-(methoxyimino)-Nmethyl-2-(2-phenoxyphenyl)acetamide, flusilazole, epoxiconazole, propiconazole and tebuconazole.
- The fungicidal composition of Claim 2 comprising a fungicidally effective 3. amount of (1) 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and (2) 2,4-dihydro-5methoxy-2-methyl-4-[2-[[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one.
- The fungicidal composition of Claim 2 comprising a fungicidally effective amount of (1) 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and (2) flusilazole.
 - The fungicidal composition of Claim 2 comprising a fungicidally effective amount of (1) 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and (2) tebuconazole.
- The fungicidal composition of Claim 2 comprising a fungicidally effective amount of (1) 6-iodo-3-propyl-2-propyloxy-4(3H)-quinazolinone and (2) both 2,4dihydro-5-methoxy-2-methyl-4-[2-[[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3H-1,2,4-triazol-3-one; and flusilazole.
- A method for controlling plant diseases caused by fungal plant pathogens 7. comprising applying to the plant or portion thereof, or to the plant seed or seedling, a fungicidally effective amount of a composition of Claim 1.
- A method of controlling at least one fungal plant disease selected from Erysiphe graminis, Puccinia recondita, Pseudocercosporella herpotrichoides and Septoria nodorum comprising applying to the plant or portion thereof, or to the plant seed or seedling
- (1) at least one compound selected from the quinazolinones of Formula I, 35 N-oxides, and agriculturally suitable salts thereof,

wherein

D is O or S;

 R^1 is C_1 - C_{10} alkyl; C_3 - C_5 cycloalkyl; C_3 - C_{10} alkenyl; C_3 - C_{10} alkynyl; 5 C_1 - C_{10} haloalkyl; C_3 - C_{10} haloalkenyl; C_3 - C_{10} haloalkynyl; C₂-C₁₀ alkoxyalkyl; C₂-C₁₀ alkylthioalkyl; C₂-C₁₀ alkylsulfinylalkyl; C2-C10 alkylsulfonylalkyl; C4-C10 cycloalkylalkyl; C_4 - C_{10} alkenyloxyalkyl; C_4 - C_{10} alkynyloxyalkyl; C_4 - C_{10} (cycloalkyl)oxyalkyl; C₄-C₁₀ alkenylthioalkyl; C₄-C₁₀ alkynylthioalkyl; 10 C₄-C₁₀ (cycloalkyl)thioalkyl; C₂-C₁₀ haloalkoxyalkyl; C₄-C₁₀ haloalkenyloxyalkyl; C₄-C₁₀ haloalkynyloxyalkyl; C₄-C₁₀ alkoxyalkenyl; C₄-C₁₀ alkoxyalkynyl; C₄-C₁₀ alkylthioalkenyl; C₄-C₁₀ alkylthioalkynyl; C₄-C₁₀ trialkylsilylalkyl; C₁-C₁₀ alkyl substituted with NR5R6, nitro, cyano or phenyl optionally substituted with R8, R9 and 15 R^{10} ; C_1 - C_{10} alkoxy; C_1 - C_{10} haloalkoxy; C_1 - C_{10} alkylthio; C₁-C₁₀ haloalkylthio; or pyridinyl, furanyl, thienyl, naphthalenyl, benzofuranyl, benzothienyl or quinolinyl each optionally substituted with R^{8} , R^{9} and R^{10} ; R^2 is C_1 - C_{10} alkyl; C_3 - C_7 cycloalkyl; C_3 - C_{10} alkenyl; C_3 - C_{10} alkynyl; 20 C₁-C₁₀ haloalkyl; C₃-C₁₀ haloalkenyl; C₃-C₁₀ haloalkynyl; C₂-C₁₀ alkoxyalkyl; C₂-C₁₀ alkylthioalkyl; C₂-C₁₀ alkylsulfinylalkyl; C2-C10 alkylsulfonylalkyl; C4-C10 cycloalkylalkyl; C_4 - C_{10} alkenyloxyalkyl; C_4 - C_{10} alkynyloxyalkyl; C_4 - C_{10} (cycloalkyl)oxyalkyl; C₄-C₁₀ alkenylthioalkyl; C₄-C₁₀ alkynylthioalkyl; 25 C₄-C₁₀ (cycloalkyl)thioalkyl; C₂-C₁₀ haloalkoxyalkyl; C₄-C₁₀ haloalkenyloxyalkyl; C₄-C₁₀ haloalkynyloxyalkyl; C₄-C₁₀ alkoxyalkenyl; C₄-C₁₀ alkoxyalkynyl; C₄-C₁₀ alkylthioalkenyl; C₄-C₁₀ alkylthioalkynyl; C₄-C₁₀ trialkylsilylalkyl; C₂-C₁₀ cyanoalkyl; C₁-C₁₀ nitroalkyl; C₁-C₁₀ alkyl substituted with CO₂R⁵, NR⁵R⁶, or phenyl 30 optionally substituted with R7, R9 and R10; phenyl optionally substituted with R^7 , R^9 and R^{10} ; -N=CR⁵R⁵; or -NR⁵R⁶; or R¹ and R² are taken together to form -CH₂(CH₂)₀CH₂-;

15

20

25

30

q is 0, 1, 2, 3 or 4;

R³ is halogen, C₁-C₈ alkyl, C₃-C₈ cycloalkyl, C₂-C₈ alkenyl, C₂-C₈ alkynyl, C₁-C₈ haloalkyl, C₃-C₈ haloalkenyl, C₃-C₈ haloalkynyl, C₁-C₈ alkoxy, C₁-C₈ haloalkoxy, C₃-C₈ alkenyloxy, C₃-C₈ alkynyloxy, C₁-C₈ alkylthio, C₃-C₈ alkenylthio, C₃-C₈ alkynylthio, C₁-C₈ alkylsulfinyl, C₁-C₈ alkylsulfinyl, C₂-C₈ alkylsulfinylalkyl, C₂-C₈ alkylsulfinylalkyl, C₄-C₈ cycloalkylalkyl, C₃-C₈ trialkylsilyl, NR⁵R⁶, C₅-C₈ trialkylsilylalkynyl or phenyl optionally substituted with at least one R⁷;

10 R⁴ is hydrogen, halogen, C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy or C₁-C₄ haloalkoxy;

each R^5 is independently hydrogen, C_1 - C_4 alkyl or phenyl optionally substituted with at least one R^7 ;

each R⁶ is independently hydrogen, C₁-C₈ alkyl or phenyl optionally substituted with at least one R⁷; or

when R⁵ and R⁶ are attached to the same nitrogen atom, R⁵ and R⁶ can be taken together to form -CH₂CH₂CH₂CH₂-, -CH₂(CH₂)₃CH₂-, -CH₂CH₂OCH₂CH₂-, -CH₂CH(Me)CH₂- or -CH₂CH(Me)OCH(Me)CH₂-;

each R^7 is independently halogen, C_1 - C_4 alkyl, C_1 - C_4 alkoxy, C_1 - C_4 haloalkyl, nitro or cyano;

 $R^8 \text{ is } C_1\text{-}C_6 \text{ alkyl, } C_1\text{-}C_6 \text{ alkoxy, } C_1\text{-}C_6 \text{ haloalkyl, halogen, } C_2\text{-}C_8 \text{ alkynyl,} \\ C_1\text{-}C_6 \text{ alkylthio, phenyl or phenoxy each optionally substituted with at least one } R^7, \text{ cyano, nitro, } C_1\text{-}C_6 \text{ haloalkoxy, } C_1\text{-}C_6 \text{ haloalkylthio, } C_2\text{-}C_6 \\ \text{alkenyl, } C_2\text{-}C_6 \text{ haloalkenyl, acetyl, } CO_2\text{Me or } N(C_1\text{-}C_2 \text{ alkyl})_2;$

each R⁹ is independently methyl, ethyl, methoxy, methylthio, halogen or trifluoromethyl; and

each R10 is independently halogen; and

- (2) at least one compound selected from (a) compounds acting at the bc_1 complex of the fungal mitochondrial respiratory electron transfer site and (b) compounds that control fungal disease by inhibiting sterol biosynthesis; wherein component (1) and component (2) are added in amounts sufficient to provide a fungicidal effectiveness greater than the sum of the fungicidal effectivenesses provided by those amounts of said components taken independently.
- 9. A method of Claim 8 wherein *Erysiphe graminis* is controlled; wherein component (1) is 6-iodo-3-propyl-2-propyloxy-4(3*H*)-quinazolinone and component (2) is selected from the group consisting of 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3*H*-1,2,4-triazol-3-one,

kresoxim-methyl, azoxystrobin, (E)-2-(methoxyimino)-N-methyl-2-(2-phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and tebuconazole; and wherein the mole ratio of component (1) to component (2) is from 4:1 to 1:10.

- 5 10. A method of Claim 8 wherein at least one of *Puccinia recondita* and *Septoria nodorum* is controlled; wherein component (1) is 6-iodo-3-propyl-2-propyloxy-4(3*H*)-quinazolinone and component (2) is selected from the group consisting of 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3*H*-1,2,4-triazol-3-one, kresoxim-methyl, azoxystrobin, (*E*)-2-(methoxyimino)-*N*-methyl-2-(2-phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and tebuconazole; and wherein the mole ratio of component (1) to component (2) is from 30:1 to 1:30.
- 11. A method of Claim 8 wherein *Pseudocercosporella herpotrichoides* is

 15 controlled; wherein component (1) is 6-iodo-3-propyl-2-propyloxy-4(3*H*)-quinazolinone and component (2) is selected from the group consisting of 2,4-dihydro-5-methoxy-2-methyl-4-[2-[[[[1-[3-(trifluoromethyl)phenyl]ethylidene]amino]oxy]methyl]phenyl]-3*H*-1,2,4-triazol-3-one, kresoxim-methyl, azoxystrobin, (*E*)-2-(methoxyimino)-*N*-methyl-2-(2-phenoxyphenyl)acetamide, flusilazole, epoxiconazole, fenpropimorph, propiconazole and tebuconazole; and wherein the mole ratio of component (1) to component (2) is from 15:1 to 1:15.

CLASSIFICATION OF SUBJECT MATTER PC 6 A01N43/54 //(A01N43/54,61:00,55:00,47:38,43:84,43:653,43:54, 37:50) According to International Patent Classification (IPC) or to both national classification and IPC Minimum documentation searched (classification system followed by classification symbols) IPC 6 A01N Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to claim No. Citation of document, with indication, where appropriate, of the relevant passages 1,2,4,5, WO 94 26722 A (DU PONT ; BEREZNAK JAMES X FRANCIS (US); CHANG ZEN YU (US); SELBY THO) 24 November 1994 cited in the application see page 1, line 10 - page 4, line 23 see page 9, line 18 - line 23 see page 33, line 3 - line 8 see page 33, line 16 - line 27 see page 33, line 33 - line 35 2,6 γ Patent family members are listed in annex. Further documents are listed in the continuation of box C. Special categories of cited documents: "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the "A" document defining the general state of the art which is not considered to be of particular relevance. "E" earlier document but published on or after the international "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to tiling date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such docu-"O" document referring to an oral disclosure, use, exhibition or ments, such combination being obvious to a person skilled in the art. other means document published prior to the international filing date but "&" document member of the same patent family later than the priority date claimed Date of malling of the international search report Date of the actual completion of theinternational search 08/06/1998 29 May 1998 Authorized officer Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijawijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016 Lamers, W

1



Inti. .tion Incation No PCT/US 98/01381

.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
ategory °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
	WO 97 02262 A (DU PONT ;BEREZNAK JAMES FRANCIS (US); CHANG ZEN YU (US); STERNBERG) 23 January 1997 see page 1, line 18 - page 3, line 30 see page 61, line 11 - line 16 see page 61, line 25 - page 62, line 4 see page 62, line 12 - page 63, line 9	1,7
	page 12, the page 12, the	2
,	WO 97 00612 A (DU PONT ;BROWN RICHARD JAMES (US); CHAN DOMINIC MING TAK (US); HOW) 9 January 1997 see page 2, line 20 - page 8, line 7 see page 117, line 6 - line 11 see page 117, line 20 - line 32	6
	see page 118, line 6 - page 119, line 6 see page 126; example 59	
٠	,	
•	,	
	·	
	·	

INTERIOR ONAL SEARCH REPORT

Inti ton Cation No PCT/US 98/01381

Patent document cited in search report		Publication date		atent family member(s)	Publication date
WO 9426722	A	24-11-1994	AU	6944394 A	12-12-1994
110 5120722	••	_,	BR	9406662 A	06-02-1996
			CA	2162846 A	24-11-1994
			CN	1123028 A	22-05-1996
			CZ	9502803 A	13-03-1996
			EP	0698013 A	28-02-1996
			HU	72829 A	28-05-1996
			JP	8510243 T	29-10-1996
			LV	11464 B	20-02-1997
		,	NZ	267222 A	28-10-1996
			PL	311641 A	04-03-1996
			SI	9420029 A	31-10-1996
			SK	140295 A	04-09-1996
			. ZA	9403112 A	06-11-1995
WO 9702262	 А	23-01-1997	AU	6478396 A	05-02-1997
no Proteot	••		EP	0836602 A	22-04-1998
WO 9700612	Α	09-01-1997	AU	6177096 A	22-01-1997
, J.	••		EP	0836384 A	22-04-1998